

=> file biosis caba caplus lifesci medline

=> e momotani e/au

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E1          1      MOMOTANDI EIKI/AU
E2          1      MOMOTANI A/AU
E3         167 --> MOMOTANI E/AU
E4          3      MOMOTANI EI ICHI/AU
E5         65      MOMOTANI EIICHI/AU
E6          2      MOMOTANI EIJI/AU
E7          8      MOMOTANI EIKI/AU
E8         27      MOMOTANI H/AU
E9         38      MOMOTANI HIROSHI/AU
E10         4      MOMOTANI HISAKO/AU
E11         1      MOMOTANI JUNICHI/AU
E12         2      MOMOTANI K/AU
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=> s e3-e7 and mycobact?

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L1          52 ("MOMOTANI E"/AU OR "MOMOTANI EI ICHI"/AU OR "MOMOTANI EIICHI"/A
              U OR "MOMOTANI EIJI"/AU OR "MOMOTANI EIKI"/AU) AND MYCOBACT?
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=> dup rem l1

PROCESSING COMPLETED FOR L1

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L2          20 DUP REM L1 (32 DUPLICATES REMOVED)
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=> d bib ab kwic 1-

YOU HAVE REQUESTED DATA FROM 20 ANSWERS - CONTINUE? Y/(N):y

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L2  ANSWER 1 OF 20  CAPLUS  COPYRIGHT 2008 ACS on STN
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AN  2007:428046  CAPLUS <<LOGINID::20080325>>
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DN  146:416306
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```
TI  Primer sets for detection of expression level of urocortin for evaluation
    of progressing of johne's disease in livestock
```

```
IN  ***Momotani, Eiichi*** ; Mori, Yasuyuki; Wang, Hong Yu
```

```
PA  National Agriculture Bio-Oriented Research Organization, Japan
```

```
SO  Jpn. Kokai Tokkyo Koho, 15pp.
```

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    CODEN: JKXXAF
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DT  Patent
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```
LA  Japanese
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FAN.CNT 1
```

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	----	-----	-----	-----
PI	JP 2007097490	A	20070419	JP 2005-291868	20051005
PRAI	JP 2005-291868		20051005		

```
AB  This invention provides primer sets for detection of expression level of
    urocortin in livestock blood sample by realtime-PCR. The cDNA sequence of
    Bos taurus urocortin were disclosed. The invention also provides method
    for prepn. of std. curve for real-time PCR by detecting the expression
    level of urocortin gene in Bos taurus cells immunized with antigen from
    ***Mycobacterium*** paratuberculosis. The method provided in this
    invention can be used for evaluation of progressing of johne's disease in
    livestock in early stage of infection.
```

```
IN  ***Momotani, Eiichi*** ; Mori, Yasuyuki; Wang, Hong Yu
```

```
AB  . . . curve for real-time PCR by detecting the expression level of
    urocortin gene in Bos taurus cells immunized with antigen from
    ***Mycobacterium*** paratuberculosis. The method provided in this
    invention can be used for evaluation of progressing of johne's disease in
    livestock in. . .
```

```
IT  ***Mycobacterium*** avium paratuberculosis
```

(infection of; primer sets for detection of expression level of urocortin for evaluation of progressing of johne's disease in livestock)

L2 ANSWER 2 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
DUPLICATE 1

AN 2007:589654 BIOSIS <<LOGINID::20080325>>

DN PREV200700590889

TI Corticotropin-releasing hormone and urocortin expression in peripheral  
blood cells from experimentally infected cattle with \*\*\*Mycobacterium\*\*\*  
avium subsp paratuberculosis.

AU Wang, Hongyu; Aodon-geril; Shu, Yujing; Momotani, Yuriko; Wang, Xiaofei;  
Mori, Yasuyuki; \*\*\*Momotani, Eiichi\*\*\* [Reprint Author]

CS Natl Inst Anim Hlth, Res Team Paratuberculosis, 3-1-5 Kan-nondai, Tsukuba,  
Ibaraki 3050856, Japan  
momotani@affrc.go.jp

SO Microbes and Infection, (JUL 2007) Vol. 9, No. 9, pp. 1061-1069.  
ISSN: 1286-4579.

DT Article

LA English

ED Entered STN: 21 Nov 2007  
Last Updated on STN: 21 Nov 2007

AB Urocortin (UCN) is a new neuropeptide of the corticotrophin-releasing  
hormone (CRH) family which plays an important role in immune responses.  
\*\*\*Mycobacterium\*\*\* avium subspecies paratuberculosis (Map) is the  
etiological agent of paratuberculosis (Johne's disease). The role of UCN  
or CRH in the pathogenesis of Map-infection is unknown. In the present  
study, we first cloned the bovine UCN gene and demonstrated the profile of  
UCN or CRH expression in peripheral blood cells from Map-infected cattle  
and uninfected controls by real-time reverse transcription-polymerase  
chain reaction (RT-PCR) and ELISA analysis. These data are the first  
observations of the characteristic kinetics of these neuropeptides in  
Map-infection. UCN or CRH expression in non-stimulated blood samples from  
infected cattle was higher than that in similarly treated samples from  
uninfected controls; however, exposure to Map lysate and live Map resulted  
in down-regulated expression of UCN in infected cattle compared to their  
counterparts from uninfected controls. These results have provided a  
direction in understanding the pathogenesis of paratuberculosis and  
improving diagnostic methods for Map-infection. (C) 2007 Elsevier Masson  
SAS. All rights reserved.

TI Corticotropin-releasing hormone and urocortin expression in peripheral  
blood cells from experimentally infected cattle with \*\*\*Mycobacterium\*\*\*  
avium subsp paratuberculosis.

AU Wang, Hongyu; Aodon-geril; Shu, Yujing; Momotani, Yuriko; Wang, Xiaofei;  
Mori, Yasuyuki; \*\*\*Momotani, Eiichi\*\*\* [Reprint Author]

AB. . . Urocortin (UCN) is a new neuropeptide of the corticotrophin-  
releasing hormone (CRH) family which plays an important role in immune  
responses. \*\*\*Mycobacterium\*\*\* avium subspecies paratuberculosis  
(Map) is the etiological agent of paratuberculosis (Johne's disease). The  
role of UCN or CRH in the. . .

ORGN . . .  
Chordata; Animalia  
Organism Name  
bovine (common): host  
Taxa Notes  
Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates,  
Nonhuman Mammals, Vertebrates

ORGN Classifier  
     \*\*\*Mycobacteriaceae\*\*\*           08881  
 Super Taxa  
     \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
     Eubacteria; Bacteria; Microorganisms  
 Organism Name  
     \*\*\*Mycobacterium\*\*\*    avium paratuberculosis (subspecies): pathogen  
 Taxa Notes  
     Bacteria, Eubacteria, Microorganisms

L2    ANSWER 3 OF 20   CABA COPYRIGHT 2008 CABI on STN  
 AN    2008:43487   CABA <<LOGINID::20080325>>  
 DN    20083029329  
 TI    Molecular strategies for studying hosts of Johne's disease  
       \*\*\*mycobacterium\*\*\*  
 AU    \*\*\*Momotani, E.\*\*\* ; Aodonqeril; Momotani, Y.  
 SO    Journal of Veterinary Medicine, Japan, (2007) Vol. 60, No. 10, pp.  
       807-813. 41 ref.  
       Publisher: Buneido Publishing Company Ltd. Tokyo  
       ISSN: 0447-0192  
       URL: <http://www.buneido-syuppan.com>

CY    Japan  
 DT    Journal  
 LA    Japanese  
 ED    Entered STN: 7 Feb 2008  
       Last Updated on STN: 7 Feb 2008  
 TI    Molecular strategies for studying hosts of Johne's disease  
       \*\*\*mycobacterium\*\*\* .  
 AU    \*\*\*Momotani, E.\*\*\* ; Aodonqeril; Momotani, Y.  
 BT    \*\*\*Mycobacterium\*\*\*   avium;   \*\*\*Mycobacterium\*\*\* ;  
       \*\*\*Mycobacteriaceae\*\*\* ; Firmicutes; bacteria; prokaryotes  
 ORGN   \*\*\*Mycobacterium\*\*\*   avium subsp. paratuberculosis

L2    ANSWER 4 OF 20   CAPLUS   COPYRIGHT 2008 ACS on STN  
 AN    2005:283672   CAPLUS <<LOGINID::20080325>>  
 DN    142:334896  
 TI    Method for diagnosing johne's disease  
 IN    \*\*\*Momotani, Eiichi\*\*\* ; Mori, Yasuyuki; Hikono, Hirokazu; Buza, Joram  
       Josephat  
 PA    Incorporated Administrative Agency National Agriculture and Bio-Oriented  
       Research Organization, Japan  
 SO    PCT Int. Appl., 38 pp.  
       CODEN: PIXXD2

DT    Patent  
 LA    Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	----	-----	-----	-----
PI	WO 2005029079	A1	20050331	WO 2003-JP11845	20030917
	W: AU, JP, US				
	RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,				
	IT, LU, MC, NL, PT, RO, SE, SI, SK, TR				
	AU 2003272880	A1	20050411	AU 2003-272880	20030917
	US 2008038758	A1	20080214	US 2007-572514	20070426
PRAI	WO 2003-JP11845	A	20030917		

AB    A method for diagnosing johne's disease is provided, with which an animal  
       infected with   \*\*\*Mycobacterium\*\*\*   paratuberculosis (Johne's) can be

diagnosed at a high sensitivity in the inapparent infection stage before the specific antibody level begins to increase, and a large no. of specimens can be treated. The method is characterized in that it comprises collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* paratuberculosis antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood. The method is also characterized in that the IFN.gamma. yield in blood is measured by the IFN.gamma. ELISA method. Also provided is a method for diagnosing \*\*\*mycobacteriosis\*\*\*, which is characterized by comprising collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood.

RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

IN \*\*\*Momotani, Eiichi\*\*\* ; Mori, Yasuyuki; Hikono, Hirokazu; Buza, Joram Josephat

AB A method for diagnosing johne's disease is provided, with which an animal infected with \*\*\*Mycobacterium\*\*\* paratuberculosis (Johne's) can be diagnosed at a high sensitivity in the inapparent infection stage before the specific antibody level begins. . . is characterized in that it comprises collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* paratuberculosis antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood. The. . . that the IFN.gamma. yield in blood is measured by the IFN.gamma. ELISA method. Also provided is a method for diagnosing \*\*\*mycobacteriosis\*\*\*, which is characterized by comprising collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood.

IT Animals

Blood analysis

Diagnosis

\*\*\*Mycobacterium\*\*\* avium paratuberculosis

(method for diagnosing johne's disease by measuring blood IFN.gamma. by ELISA)

L2 ANSWER 5 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 2

AN 2004:438665 BIOSIS <<LOGINID::20080325>>

DN PREV200400437489

TI Neutralization of interleukin-10 significantly enhances gamma interferon expression in peripheral blood by stimulation with Johnin purified protein derivative and by infection with \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in experimentally infected cattle with paratuberculosis.

AU Buza, Joram J.; Hikono, Hirokazu; Mori, Yasuyuki; Nagata, Reiko; Hirayama, Sachiyo; Bari, Abusaleh M.; Aodon-geril; Shu, Yujing; Tsuji, Noriko M.; \*\*\*Momotani, Eiichi\*\*\* [Reprint Author]

CS ParaTB and Inflammatory Bowel Dis Res Team, NIAH, 3-1-5 Kannondai, Tsukuba, Ibaraki, 3050856, Japan  
momotani@affrc.go.jp

SO Infection and Immunity, (April 2004) Vol. 72, No. 4, pp. 2425-2428. print. ISSN: 0019-9567 (ISSN print).

DT Article

LA English

ED Entered STN: 17 Nov 2004

Last Updated on STN: 17 Nov 2004

AB Monoclonal antibody neutralization of interleukin-10 (IL-10) increased Johnin purified protein derivative-induced whole-blood gamma interferon (IFN-gamma) secretion 23-fold and also increased IFN-gamma secretion ninefold following in vitro \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection of peripheral blood mononuclear cells. These results demonstrate the suppressive effect of IL-10 on immune responses to M. avium subsp. paratuberculosis infection in cattle.

TI. . . interleukin-10 significantly enhances gamma interferon expression in peripheral blood by stimulation with Johnin purified protein derivative and by infection with \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in experimentally infected cattle with paratuberculosis.

AU. . . Buza, Jorarn J.; Hikono, Hirokazu; Mori, Yasuyuki; Nagata, Reiko; Hirayama, Sachiyo; Bari, Abusaleh M.; Aodon-geril; Shu, Yujing; Tsuji, Noriko M.; \*\*\*Momotani, Eiichi\*\*\* [Reprint Author]

AB. . . increased Johnin purified protein derivative-induced whole-blood gamma interferon (IFN-gamma) secretion 23-fold and also increased IFN-gamma secretion ninefold following in vitro \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection of peripheral blood mononuclear cells. These results demonstrate the suppressive effect of IL-10 on immune responses. . .

ORGN . . .  
Animalia  
Organism Name  
cattle (common): immune responses  
Taxa Notes  
Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates,  
Nonhuman Mammals, Vertebrates

ORGN Classifier  
\*\*\*Mycobacteriaceae\*\*\* 08881  
Super Taxa  
\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
Eubacteria; Bacteria; Microorganisms  
Organism Name  
\*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies)  
Taxa Notes  
Bacteria, Eubacteria, Microorganisms

L2 ANSWER 6 OF 20 CAPLUS COPYRIGHT 2008 ACS on STN  
AN 2004:885718 CAPLUS <<LOGINID::20080325>>  
DN 141:363746  
TI Development of early-stage diagnostic method for Johne disease by using anti-IL-10 antibody  
AU \*\*\*Momotani, Eiichi\*\*\* ; Mori, Yasuyuki  
CS Natl. Agric. Bio-oriented Res. Org., Natl. Inst. Animal Health, Tsukuba, 305-0856, Japan  
SO BRAIN Techno News (2004), 105, 18-24  
CODEN: BTEEEC; ISSN: 1345-5958  
PB Nogyo, Seibutsukei Tokutei Sangyo Gijutsu Kenkyu Kiko, Seibutsukei Tokutei Sangyo Gijutsu Kenkyu Shien Senta  
DT Journal; General Review  
LA Japanese  
AB A review on early-stage diagnosis of Johne's disease (paratuberculosis) in cattle by modified interferon .gamma. ELISA assay using IL-10 neutralizing antibody, and its effectiveness.  
AU \*\*\*Momotani, Eiichi\*\*\* ; Mori, Yasuyuki  
IT Bos taurus

\*\*\*Mycobacterium\*\*\* avium paratuberculosis  
(early-stage diagnosis method for Johne's disease using anti-IL-10  
antibody)

L2 ANSWER 7 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
DUPLICATE 3  
AN 2004:64047 BIOSIS <<LOGINID::20080325>>  
DN PREV200400065534  
TI \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection causes  
suppression of RANTES, monocyte chemoattractant protein 1, and tumor  
necrosis factor alpha expression in peripheral blood of experimentally  
infected cattle.  
AU Buza, Joram J.; Mori, Yasuyuki; Bari, Abusaleh M.; Hikono, Hirokazu;  
Aodon-geril; Hirayama, Sachiyo; Shu, Yujing; \*\*\*Momotani, Eiichi\*\*\*  
[Reprint Author]  
CS Paratuberculosis and Inflammatory Bowel Disease Research Team, NIAH, 3-1-5  
Kan-nondai, Tsukuba, 305-0856, Japan  
momotani@affrc.go.jp  
SO Infection and Immunity, (December 2003) Vol. 71, No. 12, pp. 7223-7227.  
print.  
ISSN: 0019-9567 (ISSN print).  
DT Article  
LA English  
ED Entered STN: 28 Jan 2004  
Last Updated on STN: 28 Jan 2004  
AB Blood from cattle with subclinical \*\*\*Mycobacterium\*\*\* avium subsp.  
paratuberculosis infection was stimulated with M. avium subsp.  
paratuberculosis antigens, and expression of interleukin-1beta (IL-1beta),  
tumor necrosis factor alpha (TNF-alpha), RANTES, monocyte chemoattractant  
protein 1 (MCP-1), and IL-8 was measured. Expression of TNF-alpha,  
RANTES, and MCP-1 was lower in infected than in uninfected cattle. The  
reduced response may weaken protective immunity and perpetuate infection.  
TI \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection causes  
suppression of RANTES, monocyte chemoattractant protein 1, and tumor  
necrosis factor alpha expression in peripheral. . .  
AU Buza, Joram J.; Mori, Yasuyuki; Bari, Abusaleh M.; Hikono, Hirokazu;  
Aodon-geril; Hirayama, Sachiyo; Shu, Yujing; \*\*\*Momotani, Eiichi\*\*\*  
[Reprint Author]  
AB Blood from cattle with subclinical \*\*\*Mycobacterium\*\*\* avium subsp.  
paratuberculosis infection was stimulated with M. avium subsp.  
paratuberculosis antigens, and expression of interleukin-1beta (IL-1beta),  
tumor necrosis factor. . .  
ORGN . . .  
Animalia  
Organism Name  
cattle (common): host, breed-Holstein  
Taxa Notes  
Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates,  
Nonhuman Mammals, Vertebrates  
ORGN Classifier  
\*\*\*Mycobacteriaceae\*\*\* 08881  
Super Taxa  
\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
Eubacteria; Bacteria; Microorganisms  
Organism Name  
\*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen  
Taxa Notes

Bacteria, Eubacteria, Microorganisms

L2 ANSWER 8 OF 20 CAPLUS COPYRIGHT 2008 ACS on STN  
AN 2003:399194 CAPLUS <<LOGINID::20080325>>  
DN 140:39839  
TI Studies on diagnostic methods for bovine paratuberculosis  
AU Mori, Yasuyuki; Kikuma, Reiko; Muneta, Yoshihiro; Yoshihara, Kazuhiro;  
Hikono, Hirokazu; \*\*\*Momotani, Eiichi\*\*\*  
CS Immune System Section, Department of Immunology, National Institute of  
Animal Health, Tsukuba, 305-0856, Japan  
SO Dobutsu Eisei Kenkyusho Kenkyu Hokoku (2003), Volume Date 2002, 109, 33-42  
CODEN: DEKKC9; ISSN: 1347-2542  
PB Nogyo Gijutsu Kenkyu Kiko Dobutsu Eisei Kenkyusho  
DT Journal  
LA Japanese  
AB Current diagnostic tests for paratuberculosis principally rest on serol.  
assay, bacterial culture and the johnin skin test. However, diagnostic  
tests that are both sensitive and specific for detecting all subclinically  
affected animals have not yet been found. Therefore, a no. of studies  
have been conducted in order to find rapid and accurate diagnostic methods  
for paratuberculosis. As a result, the following have been found. (1)  
PCR test with internal control DNA is accurate, sensitive and rapid for  
the detection of \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in  
fecal samples. (2) In the interferon gamma (IFN-.gamma.) assay using  
johnin purified protein deriv. (J-PPD), bovine tuberculin PPD and Con A  
(Con A), IFN-.gamma. responses against J-PPD were the highest in affected  
animals. On the contrary those of Con A were the highest in healthy  
animals. Interpretation of the IFN-.gamma. assay by the higher  
IFN-.gamma. responses against J-PPD than those of Con A is preferable as  
one of the diagnostic criteria. (3) Monoclonal antibody (711-1-1) which  
recognizes the lipoarabinomannan antigen of M. avium subsp.  
paratuberculosis did not react with M. avium subsp. avium, and showed  
potential usefulness in the serol. tests. (4) A recombinant alkyl  
hydroperoxide reductase C of M. avium subsp. paratuberculosis has been  
prepd. and successfully applied to induce IFN-.gamma. from peripheral  
blood mononuclear cells of animals infected with M. avium subsp.  
paratuberculosis. (5) In the course of study on the role of cytokines,  
monocyte chemoattractant protein-1 seems to be involved in the  
pathogenesis of paratuberculosis.  
AU Mori, Yasuyuki; Kikuma, Reiko; Muneta, Yoshihiro; Yoshihara, Kazuhiro;  
Hikono, Hirokazu; \*\*\*Momotani, Eiichi\*\*\*  
AB . . . following have been found. (1) PCR test with internal control  
DNA is accurate, sensitive and rapid for the detection of  
\*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in fecal samples.  
(2)  
In the interferon gamma (IFN-.gamma.) assay using johnin purified protein  
deriv. (J-PPD), bovine. . .  
ST cattle \*\*\*Mycobacterium\*\*\* paratuberculosis infection surface antigen  
IFN induction test; alkyl hydroperoxide reductase antigen cattle IFN  
induction \*\*\*Mycobacterium\*\*\* ; reverse transcription PCR monocyte  
chemoattractant protein mRNA assay  
IT Bos taurus  
Infection  
\*\*\*Mycobacterium\*\*\* avium paratuberculosis  
(studies on diagnostic methods for bovine paratuberculosis)

L2 ANSWER 9 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN

DUPLICATE 4  
 AN 2003:329566 BIOSIS <<LOGINID::20080325>>  
 DN PREV200300329566  
 TI Studies on the diagnostic methods for bovine paratuberculosis.  
 AU Mori, Yasuyuki [Reprint Author]; Kikuma, Reiko; Muneta, Yoshihiro;  
 Yoshihara, Kazuhiro; Hikono, Hirokazu; \*\*\*Momotani, Eiichi\*\*\*  
 CS Immune System Section, Department of Immunology, National Institute of  
 Animal Health, 3-1-5 Kannondai, Tsukuba, Ibaraki, 305-0856, Japan  
 yamori@affrc.go.jp  
 SO Bulletin of the National Institute of Animal Health, (2002) No. 109, pp.  
 33-42. print.  
 ISSN: 1347-2542 (ISSN print).  
 DT Article  
 LA Japanese  
 ED Entered STN: 16 Jul 2003  
 Last Updated on STN: 16 Jul 2003  
 AB Current diagnostic tests for paratuberculosis principally rest on  
 serological assay, bacterial culture and the johnin skin test. However,  
 diagnostic tests that are both sensitive and specific for detecting all  
 subclinically affected animals have not yet been found. Therefore, a  
 number of studies have been conducted in order to find rapid and accurate  
 diagnostic methods for paratuberculosis. As a result, the following have  
 been found; 1) PCR test with internal control DNA is accurate, sensitive  
 and rapid for the detection of \*\*\*Mycobacterium\*\*\* avium subsp.  
 paratuberculosis in faecal samples. 2) In the interferon gamma (IFN-gamma)  
 assay using johnin purified protein derivative (J-PPD), bovine tuberculin  
 PPD and concanavalin A (Con A), IFN-gamma responses against J-PPD were the  
 highest in affected animals. On the contrary those of Con A were the  
 highest in healthy animals. Interpretation of the IFN-gamma assay by the  
 higher IFN-gamma responses against J-PPD than those of Con A is preferable  
 as one of the diagnostic criteria. 3) Monoclonal antibody (711-1-1) which  
 recognizes the lipoarabinomannan antigen of M. avium subsp.  
 paratuberculosis did not react with M. avium subsp. avium, and showed  
 potential usefulness in the serological tests. 4) A recombinant alkyl  
 hydroperoxide reductase C of M. avium subsp. paratuberculosis has been  
 prepared and successfully applied to induce IFN-gamma from peripheral  
 blood mononuclear cells of animals infected with M. avium subsp.  
 paratuberculosis. 5) In the course of study on the role of cytokines,  
 monocyte chemoattractant protein-1 seems to be involved in the  
 pathogenesis of paratuberculosis.  
 AU Mori, Yasuyuki [Reprint Author]; Kikuma, Reiko; Muneta, Yoshihiro;  
 Yoshihara, Kazuhiro; Hikono, Hirokazu; \*\*\*Momotani, Eiichi\*\*\*  
 AB. . . following have been found; 1) PCR test with internal control DNA is  
 accurate, sensitive and rapid for the detection of \*\*\*Mycobacterium\*\*\*  
 avium subsp. paratuberculosis in faecal samples. 2) In the interferon  
 gamma (IFN-gamma) assay using johnin purified protein derivative (J-PPD),  
 bovine. . .  
 ORGN . . .  
 Chordata; Animalia  
 Organism Name  
 bovine (common): host  
 Taxa Notes  
 Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates,  
 Nonhuman Mammals, Vertebrates  
 ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa



\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen,  
 strain-ATCC 19698, strain-Kag-1  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms

L2 ANSWER 10 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
 STN  
 AN 2000:469951 BIOSIS <<LOGINID::20080325>>  
 DN PREV200000469951  
 TI Adhesion molecules and chemokines in granulomas by \*\*\*Mycobacterium\*\*\*  
 avium subspecies paratuberculosis in TNF alpha deficient mice.  
 AU \*\*\*Momotani, E.\*\*\* ; Miyama, M.; To, T. L.; Yoshihara, K.; Gotoh, H.  
 SO Immunology Letters, (September, 2000) Vol. 73, No. 2-3, pp. 194. print.  
 Meeting Info.: 24th European Immunology Meeting of the European Federation  
 of Immunological Societies (EFIS). Poznan, Poland. September 23-26, 2000.  
 European Federation of Immunological Societies.  
 CODEN: IMLED6. ISSN: 0165-2478.  
 DT Conference; (Meeting)  
 Conference; Abstract; (Meeting Abstract)  
 LA English  
 ED Entered STN: 1 Nov 2000  
 Last Updated on STN: 10 Jan 2002  
 TI Adhesion molecules and chemokines in granulomas by \*\*\*Mycobacterium\*\*\*  
 avium subspecies paratuberculosis in TNF alpha deficient mice.  
 AU \*\*\*Momotani, E.\*\*\* ; Miyama, M.; To, T. L.; Yoshihara, K.; Gotoh, H.  
 IT Major Concepts  
 Immune System (Chemical Coordination and Homeostasis); Infection  
 IT Diseases  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis granuloma: bacterial  
 disease, clinical signs, convalescence, pathological changes  
 IT Chemicals & Biochemicals  
 RANTES: granuloma expression; fibronectin: epithelioid granuloma. . .  
 ORGN . . .  
 model, tumor necrosis factor-alpha deficiency, wild type  
 Taxa Notes  
 Animals, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals,  
 Rodents, Vertebrates  
 ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis: pathogen  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms

L2 ANSWER 11 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
 STN DUPLICATE 5  
 AN 1993:373039 BIOSIS <<LOGINID::20080325>>  
 DN PREV199396058714  
 TI Immunohistochemical distribution of S-100 alpha-positive cells in bovine  
 \*\*\*mycobacterial\*\*\* and non- \*\*\*mycobacterial\*\*\* granulomas.  
 AU \*\*\*Momotani, E.\*\*\* [Reprint author]; Kubo, M. [Reprint author];

Ishikawa, Y.; Matsubara, Y. [Reprint author]; Nakajima, Y.; Yoshino, T.  
 CS Natl. Inst. Anim. Health, 3-1-1, Kan-nondai, Tsukuba, 305, Japan  
 SO Journal of Comparative Pathology, (1993) Vol. 108, No. 3, pp. 291-301.  
 CODEN: JCVPAR. ISSN: 0021-9975.  
 DT Article  
 LA English  
 ED Entered STN: 6 Aug 1993  
 Last Updated on STN: 6 Aug 1993  
 AB By means of immunohistochemistry, the distribution of the alpha-subunit  
 (S-100-alpha) and the beta-subunit (S-100-beta) of S-100 protein was  
 studied in bovine granulomas caused by Actinomyces bovis, Actinobacillus  
 lignieresii, Actinomyces (Corynebacterium) pyogenes, Pseudomonas  
 aeruginosa, Staphylococcus aureus, \*\*\*Mycobacterium\*\*\* bovis and  
 \*\*\*Mycobacterium\*\*\* paratuberculosis. S-100-alpha-positive epithelioid  
 cells or dendritic cells were scattered among the predominantly  
 S-100-alpha-negative cells of the mononuclear phagocyte system (MPS).  
 S-100-beta was not found in the MPS cells of granulomas but was observed  
 in the endothelial cells of blood vessels. A positive reaction to S-100  
 was also seen in normal cells in the lymphoid and mammary tissues.  
 \*\*\*Mycobacterium\*\*\* granulomas contained more S-100-alpha-positive  
 cells  
 than did non- \*\*\*mycobacterial\*\*\* ones.  
 TI Immunohistochemical distribution of S-100 alpha-positive cells in bovine  
 \*\*\*mycobacterial\*\*\* and non- \*\*\*mycobacterial\*\*\* granulomas.  
 AU \*\*\*Momotani, E.\*\*\* [Reprint author]; Kubo, M. [Reprint author];  
 Ishikawa, Y.; Matsubara, Y. [Reprint author]; Nakajima, Y.; Yoshino, T.  
 AB. . . S-100 protein was studied in bovine granulomas caused by Actinomyces  
 bovis, Actinobacillus lignieresii, Actinomyces (Corynebacterium) pyogenes,  
 Pseudomonas aeruginosa, Staphylococcus aureus, \*\*\*Mycobacterium\*\*\*  
 bovis and \*\*\*Mycobacterium\*\*\* paratuberculosis. S-100-alpha-positive  
 epithelioid cells or dendritic cells were scattered among the  
 predominantly S-100-alpha-negative cells of the mononuclear phagocyte  
 system (MPS).. . . of blood vessels. A positive reaction to S-100 was  
 also seen in normal cells in the lymphoid and mammary tissues.  
 \*\*\*Mycobacterium\*\*\* granulomas contained more S-100-alpha-positive  
 cells  
 than did non- \*\*\*mycobacterial\*\*\* ones.  
 ORGN . . .  
 Classifier  
 Micrococcaceae 07702  
 Super Taxa  
 Gram-Positive Cocci; Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 Micrococcaceae  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms  
 ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms  
 ORGN Classifier  
 Pasteurellaceae 06703

Super Taxa  
Facultatively Anaerobic Gram-Negative Rods; Eubacteria; Bacteria;  
Microorganisms  
Organism. . .

L2 ANSWER 12 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
STN DUPLICATE 6  
AN 1992:369927 BIOSIS <<LOGINID::20080325>>  
DN PREV199294051977; BA94:51977  
TI IMMUNOHISTOCHEMICAL IDENTIFICATION OF FERRITIN LACTOFERRIN AND TRANSFERRIN  
IN LEPROSY LESIONS OF HUMAN SKIN BIOPSIES.  
AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; WUSCHER N; RAVISSE P; RASTOGI N  
CS LAB IMMUNOPATHOL, NATIONAL INST ANIMAL HEALTH, 3-1-1 KANNONDAI, TSUKUBA,  
IBARAKI 305, JPN  
SO Journal of Comparative Pathology, (1992) Vol. 106, No. 3, pp. 213-220.  
CODEN: JCVPAR. ISSN: 0021-9975.  
DT Article  
FS BA  
LA ENGLISH  
ED Entered STN: 9 Aug 1992  
Last Updated on STN: 9 Aug 1992  
AB Granulomatous lesions of human leprosy contained ferritin and lactoferrin  
but little or no transferrin, as demonstrated by the avidin-biotin complex  
immunoperoxidase method. Lactoferrin was found in the neutrophils. These  
results suggested that the cells of the host mononuclear phagocyte system  
in leprosy granulomas provide an adequate nutritional environment for iron  
acquisition by \*\*\*Mycobacterium\*\*\* leprae. A possible role of iron  
binding proteins in the granulomas is discussed in relation to previous  
data on bovine paratuberculous granulomas.  
AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; WUSCHER N; RAVISSE P; RASTOGI N  
AB. . . the cells of the host mononuclear phagocyte system in leprosy  
granulomas provide an adequate nutritional environment for iron  
acquisition by \*\*\*Mycobacterium\*\*\* leprae. A possible role of iron  
binding proteins in the granulomas is discussed in relation to previous  
data on bovine. . .  
IT Miscellaneous Descriptors  
\*\*\*MYCOBACTERIUM\*\*\* -LEPRAE NEUTROPHILS IRON BINDING PROTEINS  
ORGN Classifier  
\*\*\*Mycobacteriaceae\*\*\* 08881  
Super Taxa  
\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
Eubacteria; Bacteria; Microorganisms  
Taxa Notes  
Bacteria, Eubacteria, Microorganisms  
ORGN Classifier  
Hominidae 86215  
Super Taxa  
Primates; Mammalia;. . .

L2 ANSWER 13 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
STN DUPLICATE 7  
AN 1991:182263 BIOSIS <<LOGINID::20080325>>  
DN PREV199191097012; BA91:97012  
TI IMMUNOHISTOCHEMICAL STUDY OF BOVINE LYMPH NODES WITH ANTIBODIES AGAINST  
S100 PROTEIN SUBUNITS COMPARISON BETWEEN LYMPH NODES OF HEALTHY AND  
\*\*\*MYCOBACTERIUM\*\*\* -PARATUBERCULOSIS INFECTED CATTLE.  
AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; YOSHINO T; ISHIKAWA Y; NAKAJIMA Y

CS HOKKAIDO BRANCH LAB, NATL INST ANIM HEALTH, HITSUJIGAOKA-4, TOYOHIRA,  
SAPPORO 004, JPN

SO Research in Immunology, (1990) Vol. 141, No. 8, pp. 771-782.  
CODEN: RIMME5. ISSN: 0923-2494.

DT Article

FS BA

LA ENGLISH

ED Entered STN: 19 Apr 1991  
Last Updated on STN: 20 Apr 1991

AB Using immunohistochemistry, the differential distribution of the .alpha.  
subunit (S100.alpha.) and .beta. subunit (S100.beta.) of S100 protein was  
studied in mesenteric lymph nodes from normal or \*\*\*Mycobacterium\*\*\*  
paratuberculosis-infected cattle. In epithelioid cell granulomas,  
S100.alpha.-positive epithelioid cells and some giant cells were scattered  
among S100.alpha.-negative cells, which were predominant. The  
S100.beta.-positive and -negative cells contained acid-fast bacilli. The  
presence of S100.beta.-positive cells was not demonstrated in the  
granulomas. In normal component cells in the lymph nodes, follicular  
dendritic cells in the germinal centres and endothelium of lymphatic sinus  
and lymph vessels were positive for S100.alpha.. S100.beta. was positive  
only in the endothelial cells of blood vessels. Results shown in the  
present paper are discussed in light of results obtained in other work on  
human tissues using the same sources of antibodies.

TI IMMUNOHISTOCHEMICAL STUDY OF BOVINE LYMPH NODES WITH ANTIBODIES AGAINST  
S100 PROTEIN SUBUNITS COMPARISON BETWEEN LYMPH NODES OF HEALTHY AND  
\*\*\*MYCOBACTERIUM\*\*\* -PARATUBERCULOSIS INFECTED CATTLE.

AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; YOSHINO T; ISHIKAWA Y; NAKAJIMA Y

AB. . . the .alpha. subunit (S100.alpha.) and .beta. subunit (S100.beta.) of  
S100 protein was studied in mesenteric lymph nodes from normal or  
\*\*\*Mycobacterium\*\*\* paratuberculosis-infected cattle. In epithelioid  
cell granulomas, S100.alpha.-positive epithelioid cells and some giant  
cells were scattered among S100.alpha.-negative cells, which were. . .

ORGN Classifier  
\*\*\*Mycobacteriaceae\*\*\* 08881  
Super Taxa  
\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
Eubacteria; Bacteria; Microorganisms  
Taxa Notes  
Bacteria, Eubacteria, Microorganisms

ORGN Classifier  
Bovidae 85715  
Super Taxa  
Artiodactyla; Mammalia;. . .

L2 ANSWER 14 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
STN DUPLICATE 8

AN 1989:223916 BIOSIS <<LOGINID::20080325>>

DN PREV198987115533; BA87:115533

TI IMMUNOHISTOCHEMICAL LOCALIZATION OF IMMUNOGLOBULINS IN BOVINE  
GRANULOMATOUS LESIONS.

AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; KUBO M; ISHIKAWA Y; YOSHINO T

CS HOKKAIDO BRANCH LAB, NATL INST ANIM HEALTH, HITSUJIGAOKA 4, TOYOHIRA,  
SAPPORO 004 JPN

SO Journal of Comparative Pathology, (1989) Vol. 100, No. 2, pp. 129-136.  
CODEN: JCVPAR. ISSN: 0021-9975.

DT Article

FS BA

LA ENGLISH  
 ED Entered STN: 7 May 1989  
 Last Updated on STN: 7 May 1989  
 AB The immunohistochemical distribution of IgG, IgA and IgM in granulomatous lesions caused by Actinomyces bovis, Actinobacillus lignieresii, Actinomyces (Corynebacterium) pyogenes, Pseudomonas aeruginosa, Staphylococcus aureus and \*\*\*Mycobacterium\*\*\* bovis was studied. Numerous IgG-containing cells (plasma cells) were distributed in the peripheral connective tissue layers, but not in the epithelioid cell layer. A few scattered IgA- and IgM-containing cells were found in all the lesions examined. \*\*\*Mycobacterial\*\*\* granulomas contained fewer IgG-cells than did non- \*\*\*mycobacterial\*\*\* granulomas. Eosinophilic club-shaped bodies were found in A. bovis, A. lignieresii, P. aeruginosa and S. aureus, but they were generally negative for IgG, IgA and IgM.  
 AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; KUBO M; ISHIKAWA Y; YOSHINO T  
 AB. . . IgA and IgM in granulomatous lesions caused by Actinomyces bovis, Actinobacillus lignieresii, Actinomyces (Corynebacterium) pyogenes, Pseudomonas aeruginosa, Staphylococcus aureus and \*\*\*Mycobacterium\*\*\* bovis was studied. Numerous IgG-containing cells (plasma cells) were distributed in the peripheral connective tissue layers, but not in the epithelioid cell layer. A few scattered IgA- and IgM-containing cells were found in all the lesions examined. \*\*\*Mycobacterial\*\*\* granulomas contained fewer IgG-cells than did non- \*\*\*mycobacterial\*\*\* granulomas. Eosinophilic club-shaped bodies were found in A. bovis, A. lignieresii, P. aeruginosa and S. aureus, but they were generally. . .  
 IT Miscellaneous Descriptors  
 ACTINOMYCES-BOVIS ACTINOBACILLUS-LIGNIERESII ACTINOMYCES-PYOGENES  
 PSEUDOMONAS-AERUGINOSA STAPHYLOCOCCUS-AUREUS \*\*\*MYCOBACTERIUM\*\*\*  
 -BOVIS IMMUNOGLOBULIN G IMMUNOGLOBULIN A IMMUNOGLOBULIN M PERIPHERAL  
 CONNECTIVE TISSUE  
 ORGN . . .  
 Nonsporing Gram-Positive Rods 08890  
 Super Taxa  
 Actinomycetes and Related Organisms; Eubacteria; Bacteria;  
 Microorganisms  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms  
 ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms  
 ORGN Classifier  
 Bovidae 85715  
 Super Taxa  
 Artiodactyla; Mammalia;. . .  
 L2 ANSWER 15 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
 STN DUPLICATE 9  
 AN 1988:483621 BIOSIS <<LOGINID::20080325>>  
 DN PREV198886114931; BA86:114931  
 TI THE DISTRIBUTION OF FERRITIN LACTOFERRIN AND TRANSFERRIN IN GRANULOMATOUS LYMPHADENITIS OF BOVINE PARATUBERCULOSIS.  
 AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; WHIPPLE D L; THIERMANN A B  
 CS HOKKAIDO BRANCH LAB, NATL INST ANIM HEALTH, HITSUJIGAOKA-4, SAPPORO 004

JPN

SO Journal of Comparative Pathology, (1988) Vol. 99, No. 2, pp. 205-214.  
CODEN: JCVPAR. ISSN: 0021-9975.

DT Article

FS BA

LA ENGLISH

ED Entered STN: 1 Nov 1988  
Last Updated on STN: 1 Nov 1988

AB Immunohistochemical examination of iron-binding proteins was carried out in the formalin-fixed mesenteric lymph nodes of normal cattle and of cattle with paratuberculosis. Ferritin (FT) and lactoferrin (LF) were found in the granulomas in ileal lymph nodes from six infected cattle. A weak reaction for transferrin (TF) was found in granulomas of a lymph node from one of the infected cattle. FT was found in the macrophages in the medullary sinuses of normal and infected nodes; however, the reaction in infected nodes was generally stronger than that in normal ones. LF in the macrophages was found in only two infected nodes. Neutrophils in both normal and infected cattle always reacted strongly for LF. The TF was always found in the blood vessels and intracellular space. These results suggest that: (1) FT and LF may be important in vivo sources of iron for \*\*\*Mycobacterium\*\*\* paratuberculosis, since their own iron-binding compounds are considered to acquire iron from FT and LF in vitro; (2) the increase in FT and LF in the granulomas may be related to inflammatory hyposideraemia associated with paratuberculosis and (3) epithelioid and giant cells may have a different iron metabolism, from normal macrophages.

AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; WHIPPLE D L; THIERMANN A B

AB. . . and intracellular space. These results suggest that: (1) FT and LF may be important in vivo sources of iron for \*\*\*Mycobacterium\*\*\* paratuberculosis, since their own iron-binding compounds are considered to acquire iron from FT and LF in vitro; (2) the increase. . .

IT Miscellaneous Descriptors  
CATTLE \*\*\*MYCOBACTERIUM\*\*\* -PARATUBERCULOSIS MACROPHAGE NEUTROPHIL  
HYPOSIDEREMIA

ORGN Classifier  
\*\*\*Mycobacteriaceae\*\*\* 08881  
Super Taxa  
\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
Eubacteria; Bacteria; Microorganisms  
Taxa Notes  
Bacteria, Eubacteria, Microorganisms

ORGN Classifier  
Bovidae 85715  
Super Taxa  
Artiodactyla; Mammalia;. . .

L2 ANSWER 16 OF 20 CABA COPYRIGHT 2008 CABI on STN DUPLICATE 10

AN 88:65054 CABA <<LOGINID::20080325>>

DN 19882208990

TI Role of M cells and macrophages in the entrance of \*\*\*Mycobacterium\*\*\* paratuberculosis into domes of ileal Peyer's patches in calves

AU \*\*\*Momotani, E.\*\*\* ; Whipple, D. L.; Thiermann, A. B.; Cheville, N. F.

CS Nat. Anim. Dis. Center, PO Box 70, Ames, IA 50010, USA.

SO Veterinary Pathology, (1988) Vol. 25, No. 2, pp. 131-137. 20 ref.  
ISSN: 0300-9858

DT Journal

LA English

ED Entered STN: 1 Nov 1994

Last Updated on STN: 1 Nov 1994

AB Ligated ileal loops of calves inoculated with live and heat-killed *M. paratuberculosis* were examined by light and electron microscopy. At 5 hours after inoculation, acid-fast bacilli were in subepithelial macrophages, but not in M cells covering domes. At 20 hours, more than 50 acid-fast bacilli per cross section were in subepithelial macrophages in domes. Both living and heat-killed bacilli passed into domes. Addition of anti-*M. paratuberculosis* bovine serum to the inoculum increased entry of bacteria into domes. Electron microscopy showed intact bacilli with electron-transparent zones (peribacillary spaces) in the supranuclear cytoplasm of M cells at 20 hours. M cells also contained vacuoles, including electron-dense material interpreted as degraded bacilli. Subepithelial and intraepithelial macrophages contained bacilli and degraded bacterial material in phagosomes. These results suggest that calf ileal M cells take up bacilli, and that subepithelial and intraepithelial macrophages secondarily accept bacilli or bacterial debris which are expelled from M cells.

TI Role of M cells and macrophages in the entrance of \*\*\**Mycobacterium*\*\*\* paratuberculosis into domes of ileal Peyer's patches in calves.

AU \*\*\*Momotani, E.\*\*\* ; Whipple, D. L.; Thiermann, A. B.; Cheville, N. F.

BT mammals; vertebrates; Chordata; animals; young animals;

\*\*\**Mycobacterium*\*\*\* ; \*\*\**Mycobacteriaceae*\*\*\* ; Firmicutes; bacteria; prokaryotes

ORGN \*\*\**Mycobacterium*\*\*\* paratuberculosis; cattle

L2 ANSWER 17 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
STN DUPLICATE 11

AN 1987:106473 BIOSIS <<LOGINID::20080325>>

DN PREV198783055451; BA83:55451

TI IMMUNOHISTOCHEMICAL DISTRIBUTION OF IMMUNOGLOBULIN AND SECRETORY COMPONENT IN THE ILEUM OF NORMAL AND PARATUBERCULOSIS-INFECTED CATTLE.

AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; ISHIKAWA Y; YOSHINO T

CS HOKKAIDO BRANCH LABORATORY, NATIONAL INSTITUTE OF ANIMAL HEALTH,  
HITSUJIGAOKA 4, TOYOHIRA-KU, SAPPORO 004, JAPAN

SO Journal of Comparative Pathology, (1986) Vol. 96, No. 6, pp. 659-670.  
CODEN: JCVPAR. ISSN: 0021-9975.

DT Article

FS BA

LA ENGLISH

ED Entered STN: 26 Feb 1987

Last Updated on STN: 26 Feb 1987

AB The immunohistochemical distribution of IgA, IgG, IgM and secretory component in the ileum of 10 normal and 21 paratuberculosis-infected cattle was investigated. Semi-quantitative analysis of the number of each class of Ig-containing cells in the lamina propria mucosa of infected ileums showed that IgG and IgM-containing cells and total Ig-containing cells were significantly more numerous than those in the normal ileums. There was no significant difference in the numbers of IgA-containing cells between the two groups of cattle. The distribution of IgA, IgM and SC was basically similar in the two groups. However, IgG-containing cells characteristically accumulated around the granulomas. It was considered that excessive local production of Ig in the intestinal mucosa, along with subsequent formation of immune complex or release of histamine from mast cells, could account for the occurrence of diarrhoea and participate in the pathogenesis of bovine paratuberculosis. A comparison of the local immunological state in paratuberculosis and Crohn's disease was made.

AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; ISHIKAWA Y; YOSHINO T

```

ORGN Classifier
      ***Mycobacteriaceae***      08881
Super Taxa
      ***Mycobacteria*** ; Actinomycetes and Related Organisms;
      Eubacteria; Bacteria; Microorganisms
Taxa Notes
      Bacteria, Eubacteria, Microorganisms
ORGN Classifier
      Bovidae      85715
Super Taxa
      Artiodactyla; Mammalia;. . .

L2  ANSWER 18 OF 20  BIOSIS  COPYRIGHT (c) 2008 The Thomson Corporation  on
STN                                     DUPLICATE 12
AN  1986:281052  BIOSIS <<LOGINID::20080325>>
DN  PREV198682024915; BA82:24915
TI  IMMUNOHISTOCHEMICAL DISTRIBUTION OF FERRITIN LACTOFERRIN AND TRANSFERRIN
    IN GRANULOMAS OF BOVINE PARATUBERCULOSIS.
AU  ***MOMOTANI E*** [Reprint author]; FURUGOURI K; OBARA Y; MIYATA Y;
    ISHIKAWA Y; YOSHINO T
CS  HAKKAIDO BRANCH LABORATORY, NATIONAL INSTITUTE OF ANIMAL HEALTH,
    HITSUJIGAOKA, TOYOHIRA-KU, SAPPORO 004, JAPAN
SO  Infection and Immunity, (1986) Vol. 52, No. 2, pp. 623-627.
    CODEN: INFIBR. ISSN: 0019-9567.
DT  Article
FS  BA
LA  ENGLISH
ED  Entered STN: 4 Jul 1986
    Last Updated on STN: 4 Jul 1986
AB  Granulomatous lesions of bovine paratuberculosis contained ferritin,
    lactoferrin and a small amount of transferrin, as demonstrated by the
    immunohistochemical method. Macrophages in the normal bovine ileum did
    not contain lactoferrin and transferrin; however, ferritin was found in
    individual macrophages of Peyer's patches. These results may help
    elucidate the relationship between intracellular growth of
    ***Mycobacterium*** paratuberculosis and the presence of iron-binding
    proteins in the granulomas.
AU  ***MOMOTANI E*** [Reprint author]; FURUGOURI K; OBARA Y; MIYATA Y;
    ISHIKAWA Y; YOSHINO T
AB. . . ferritin was found in individual macrophages of Peyer's patches.
    These results may help elucidate the relationship between intracellular
    growth of ***Mycobacterium*** paratuberculosis and the presence of
    iron-binding proteins in the granulomas.
IT  Miscellaneous Descriptors
    ***MYCOBACTERIUM*** -PARATUBERCULOSIS      ***MYCOBACTIN***
EXOCHELIN
    IRON-BINDING PROTEIN INTRACELLULAR GROWTH ENTERITIS
ORGN Classifier
      ***Mycobacteriaceae***      08881
Super Taxa
      ***Mycobacteria*** ; Actinomycetes and Related Organisms;
      Eubacteria; Bacteria; Microorganisms
Taxa Notes
      Bacteria, Eubacteria, Microorganisms
ORGN Classifier
      Bovidae      85715
Super Taxa

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Artiodactyla; Mammalia;. . .  
RN 1400-46-0 ( \*\*\*MYCOBACTIN\*\*\* )

L2 ANSWER 19 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
STN DUPLICATE 13

AN 1985:408087 BIOSIS <<LOGINID::20080325>>  
DN PREV198580078079; BA80:78079  
TI CASEOUS GRANULOMAS IN BOVINE PARATUBERCULOSIS.  
AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; YOSHINO T  
CS HODDAIDO BRANCH LABORATORY, NATIONAL INSTITUTE OF ANIMAL HEALTH, 4  
HITSUJIGAOKA, TOYOHIRA-KU, SAPPORO 061-01, JAPAN  
SO Japanese Journal of Veterinary Science, (1985) Vol. 47, No. 3, pp.  
487-492.  
CODEN: NJUZA9. ISSN: 0021-5295.

DT Article  
FS BA  
LA ENGLISH  
AB Caseous granulomas were found in the mesenteric lymph nodes of an  
Aberdeen-Angus cow, 5 yr old, having paratuberculosis. They were found  
together with epithelioid cell granulomas, and both of them contained  
numerous acid-fast bacilli. The intestine showed extensive  
paratuberculous lesions with numerous acid-fast bacilli. There was no  
caseation but focal neutrophil infiltration in the intestinal granulomas.  
\*\*\*Mycobacterium\*\*\* paratuberculosis was isolated from both the  
mesenteric lymph nodes and the intestine. No tuberculous bacilli were  
detected, and tuberculin and Johnin tests were negative.

AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; YOSHINO T  
AB. . . showed extensive paratuberculous lesions with numerous acid-fast  
bacilli. There was no caseation but focal neutrophil infiltration in the  
intestinal granulomas. \*\*\*Mycobacterium\*\*\* paratuberculosis was  
isolated from both the mesenteric lymph nodes and the intestine. No  
tuberculous bacilli were detected, and tuberculin and. . .

IT Miscellaneous Descriptors  
\*\*\*MYCOBACTERIUM\*\*\* -PARATUBERCULOSIS INTESTINAL PARATUBERCULOUS  
LESIONS ACID-FAST BACILLI FOCAL NEUTROPHIL INFILTRATION

ORGN Classifier  
\*\*\*Mycobacteriaceae\*\*\* 08881  
Super Taxa  
\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
Eubacteria; Bacteria; Microorganisms  
Taxa Notes  
Bacteria, Eubacteria, Microorganisms

ORGN Classifier  
Bovidae 85715  
Super Taxa  
Artiodactyla; Mammalia;. . .

L2 ANSWER 20 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
STN DUPLICATE 14

AN 1985:279500 BIOSIS <<LOGINID::20080325>>  
DN PREV198579059496; BA79:59496  
TI PATHOLOGICAL CHANGES OF SPONTANEOUS DUAL INFECTION OF TUBERCULOSIS AND  
PARATUBERCULOSIS IN BEEF CATTLE.  
AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; YOSHINO T  
CS HOKKAIDO BRANCH LAB, NATL INST ANIMAL HEALTH, 4 HITSUJIGAOKA, TOYOHIRA-KU,  
SAPPORO, HOKKAIDO 061-01, JPN  
SO Japanese Journal of Veterinary Science, (1984) Vol. 46, No. 5, pp.

625-632.  
 CODEN: NJUZA9. ISSN: 0021-5295.

DT Article  
 FS BA  
 LA ENGLISH  
 AB Four cases of spontaneous dual infection of tuberculosis and paratuberculosis in beef cattle, first noticed in Japan, were examined pathologically. All of the cattle were from the same limited area of Hokkaido island. Tuberculous lesions were found in 2-9 organs, including the liver, lung, kidney and mesenteric lymph nodes, and granulomas showed the same appearance as in the case of single infection. Paratuberculous lesions characterized by intracellular short acid-fast bacilli were found in the jejunum, ileum, cecum and the draining lymph nodes, but they were not severe. The mesenteric lymph nodes revealed both tuberculous and paratuberculous lesions, and in 2 of the 4 cases both types of granulomas were present in the same sections. In such sections, short acid-fast bacilli were not numerous in granulomas adjacent to tuberculous lesions.

AU \*\*\*MOMOTANI E\*\*\* [Reprint author]; YOSHINO T  
 ORGN Classifier  
     \*\*\*Mycobacteriaceae\*\*\* 08881  
     Super Taxa  
         \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
         Eubacteria; Bacteria; Microorganisms  
     Taxa Notes  
         Bacteria, Eubacteria, Microorganisms

ORGN Classifier  
     Bovidae 85715  
     Super Taxa  
         Artiodactyla; Mammalia;. . .

=> e mori yasuyuki/au

E1	58	MORI YASUYOSHI/AU
E2	1	MORI YASUYOSMI/AU
E3	175 -->	MORI YASUYUKI/AU
E4	1	MORI YASUZANE/AU
E5	12	MORI YAYOI/AU
E6	233	MORI YO/AU
E7	1	MORI YO ICHI/AU
E8	1	MORI YOHIRO/AU
E9	4	MORI YOHKO/AU
E10	3	MORI YOHTA/AU
E11	311	MORI YOICHI/AU
E12	56	MORI YOICHIRO/AU

=> s e3 and mycobact?

L3 33 "MORI YASUYUKI"/AU AND MYCOBACT?

=> dup rem l3

PROCESSING COMPLETED FOR L3

L4 15 DUP REM L3 (18 DUPLICATES REMOVED)

=> d bib ab kwic 1-

YOU HAVE REQUESTED DATA FROM 15 ANSWERS - CONTINUE? Y/(N):y

L4 ANSWER 1 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN  
 AN 2007:428046 CAPLUS <<LOGINID::20080325>>

DN 146:416306  
TI Primer sets for detection of expression level of urocortin for evaluation  
of progressing of johne's disease in livestock  
IN Momotani, Eiichi; \*\*\*Mori, Yasuyuki\*\*\* ; Wang, Hong Yu  
PA National Agriculture Bio-Oriented Research Organization, Japan  
SO Jpn. Kokai Tokkyo Koho, 15pp.  
CODEN: JKXXAF

DT Patent  
LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	----	-----	-----	-----
PI	JP 2007097490	A	20070419	JP 2005-291868	20051005
PRAI	JP 2005-291868		20051005		

AB This invention provides primer sets for detection of expression level of urocortin in livestock blood sample by realtime-PCR. The cDNA sequence of Bos taurus urocortin were disclosed. The invention also provides method for prepn. of std. curve for real-time PCR by detecting the expression level of urocortin gene in Bos taurus cells immunized with antigen from \*\*\*Mycobacterium\*\*\* paratuberculosis. The method provided in this invention can be used for evaluation of progressing of johne's disease in livestock in early stage of infection.

IN Momotani, Eiichi; \*\*\*Mori, Yasuyuki\*\*\* ; Wang, Hong Yu

AB . . . curve for real-time PCR by detecting the expression level of urocortin gene in Bos taurus cells immunized with antigen from \*\*\*Mycobacterium\*\*\* paratuberculosis. The method provided in this invention can be used for evaluation of progressing of johne's disease in livestock in. . .

IT \*\*\*Mycobacterium\*\*\* avium paratuberculosis  
(infection of; primer sets for detection of expression level of urocortin for evaluation of progressing of johne's disease in livestock)

L4 ANSWER 2 OF 15 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
DUPLICATE 1

AN 2007:589654 BIOSIS <<LOGINID::20080325>>

DN PREV200700590889

TI Corticotropin-releasing hormone and urocortin expression in peripheral blood cells from experimentally infected cattle with \*\*\*Mycobacterium\*\*\* avium subsp paratuberculosis.

AU Wang, Hongyu; Aodon-geril; Shu, Yujing; Momotani, Yuriko; Wang, Xiaofei;  
\*\*\*Mori, Yasuyuki\*\*\* ; Momotani, Eiichi [Reprint Author]

CS Natl Inst Anim Hlth, Res Team Paratuberculosis, 3-1-5 Kan-nondai, Tsukuba, Ibaraki 3050856, Japan  
momotani@affrc.go.jp

SO Microbes and Infection, (JUL 2007) Vol. 9, No. 9, pp. 1061-1069.  
ISSN: 1286-4579.

DT Article

LA English

ED Entered STN: 21 Nov 2007

Last Updated on STN: 21 Nov 2007

AB Urocortin (UCN) is a new neuropeptide of the corticotrophin-releasing hormone (CRH) family which plays an important role in immune responses. \*\*\*Mycobacterium\*\*\* avium subspecies paratuberculosis (Map) is the etiological agent of paratuberculosis (Johne's disease). The role of UCN or CRH in the pathogenesis of Map-infection is unknown. In the present study, we first cloned the bovine UCN gene and demonstrated the profile of

UCN or CRH expression in peripheral blood cells from Map-infected cattle and uninfected controls by real-time reverse transcription-polymerase chain reaction (RT-PCR) and ELISA analysis. These data are the first observations of the characteristic kinetics of these neuropeptides in Map-infection. UCN or CRH expression in non-stimulated blood samples from infected cattle was higher than that in similarly treated samples from uninfected controls; however, exposure to Map lysate and live Map resulted in down-regulated expression of UCN in infected cattle compared to their counterparts from uninfected controls. These results have provided a direction in understanding the pathogenesis of paratuberculosis and improving diagnostic methods for Map-infection. (C) 2007 Elsevier Masson SAS. All rights reserved.

TI Corticotropin-releasing hormone and urocortin expression in peripheral blood cells from experimentally infected cattle with \*\*\*Mycobacterium\*\*\* avium subsp paratuberculosis.

AU Wang, Hongyu; Aodon-geril; Shu, Yujing; Momotani, Yuriko; Wang, Xiaofei; \*\*\*Mori, Yasuyuki\*\*\* ; Momotani, Eiichi [Reprint Author]

AB. . . Urocortin (UCN) is a new neuropeptide of the corticotrophin-releasing hormone (CRH) family which plays an important role in immune responses. \*\*\*Mycobacterium\*\*\* avium subspecies paratuberculosis (Map) is the etiological agent of paratuberculosis (Johne's disease). The role of UCN or CRH in the. . .

ORGN . . .  
 Chordata; Animalia  
 Organism Name  
 bovine (common): host  
 Taxa Notes  
 Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Vertebrates

ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms

L4 ANSWER 3 OF 15 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 2

AN 2008:30137 BIOSIS <<LOGINID::20080325>>

DN PREV200800031655

TI Detection of \*\*\*Mycobacterium\*\*\* avium subsp paratuberculosis in ovine faeces by direct quantitative PCR has similar or greater sensitivity compared to radiometric culture.

AU Kawaji, Satoko; Taylor, Deborah L.; \*\*\*Mori, Yasuyuki\*\*\* ; Whittington, Richard J. [Reprint Author]

CS Univ Sydney, Fac Vet Sci, 425 Werombi Rd, Camden, NSW 2570, Australia richardw@camden.usyd.edu.au

SO Veterinary Microbiology, (NOV 15 2007) Vol. 125, No. 1-2, pp. 36-48. CODEN: VMICDQ. ISSN: 0378-1135.

DT Article

LA English

ED Entered STN: 19 Dec 2007  
 Last Updated on STN: 19 Dec 2007

AB The aims of this study were to develop a new real-time quantitative PCR

(QPCR) assay based on IS900 for detection and quantification of  
 \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis (MAP) DNA in faeces,  
 and to use this to detect infected sheep. Both the C and S strains of MAP  
 were detected by the QPCR assay, and no cross reactions were detected with  
 51 other species of \*\*\*mycobacteria\*\*\* including 10 which contained  
 IS900-like sequences. One copy of IS900 fragment cloned into plasmid  
 pCR2.1 and 1 fg of MAP genomic DNA were consistently detected, while in  
 spiked faecal samples the detection limit was 10 viable MAP per gram of  
 ovine faeces. A total of 506 individual ovine faecal samples and 27  
 pooled ovine faecal samples with known culture results were tested. The  
 QPCR assay detected 68 of 69 BACTEC culture positive individual faeces and  
 there was a strong relation between time to detection in culture and DNA  
 quantity measured by QPCR ( $r = -0.70$ ). In pooled faecal samples, QPCR  
 also agreed with culture ( $\kappa = 0.59$ ). MAP DNA was detected from some  
 culture negative faecal samples from sheep exposed to MAP, suggesting that  
 the QPCR has very high analytical sensitivity for MAP in faecal samples  
 and detects non-viable MAP in ovine faeces. None of the faecal samples  
 from 176 sheep that were not exposed to MAP were positive in QPCR. This  
 is the first report of a direct faecal QPCR assay that has similar  
 sensitivity to a gold standard radiometric culture assay. (C) 2007  
 Elsevier B.V. All rights reserved.

TI Detection of \*\*\*Mycobacterium\*\*\* avium subsp paratuberculosis in ovine  
 faeces by direct quantitative PCR has similar or greater sensitivity  
 compared to radiometric culture.

AU Kawaji, Satoko; Taylor, Deborah L.; \*\*\*Mori, Yasuyuki\*\*\* ; Whittington,  
 Richard J. [Reprint Author]

AB. . . this study were to develop a new real-time quantitative PCR (QPCR)  
 assay based on IS900 for detection and quantification of  
 \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis (MAP) DNA in faeces,  
 and to use this to detect infected sheep. Both the C and S. . .  
 strains of MAP were detected by the QPCR assay, and no cross reactions  
 were detected with 51 other species of \*\*\*mycobacteria\*\*\* including 10  
 which contained IS900-like sequences. One copy of IS900 fragment cloned  
 into plasmid pCR2.1 and 1 fg of MAP. . .

ORGN . . .  
 Chordata; Animalia  
 Organism Name  
 ovine (common): host  
 Taxa Notes  
 Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates,  
 Nonhuman Mammals, Vertebrates

ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms

L4 ANSWER 4 OF 15 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
 DUPLICATE 3

AN 2006:532033 BIOSIS <<LOGINID::20080325>>

DN PREV200600524060

TI A highly sensitive and subspecies-specific surface antigen enzyme-linked  
 immunosorbent assay for diagnosis of Johne's disease.

AU   Eda, Shigetoshi; Bannantine, John P.; Waters, W. R.;   \*\*\*Mori,\*\*\*  
 \*\*\*       Yasuyuki\*\*\*   ; Whitlock, Robert H.; Scott, M. Cathy; Speer, C. A.  
 [Reprint  
   Author]  
 CS   Univ Tennessee, Ctr Wildlife Hlth, Dept Forestry Wildlife and Fisheries,  
      POB 1071, Knoxville, TN 37901 USA  
      caspeer@utk.edu  
 SO   Clinical and Vaccine Immunology, (AUG 2006) Vol. 13, No. 8, pp. 837-844.  
      ISSN: 1556-6811.  
 DT   Article  
 LA   English  
 ED   Entered STN: 12 Oct 2006  
      Last Updated on STN: 12 Oct 2006  
 AB   Johne's disease (JD), or paratuberculosis, caused by   \*\*\*Mycobacterium\*\*\*  
      avium subsp. paratuberculosis, is one of the most widespread and  
      economically important diseases of livestock and wild ruminants worldwide.  
      Control of JD could be accomplished by diagnosis and good animal  
      husbandry, but this is currently not feasible because commercially  
      available diagnostic tests have low sensitivity levels and are incapable  
      of diagnosing prepatent infections. In this study, a highly sensitive and  
      subspecies-specific enzyme-linked immunosorbent assay was developed for  
      the diagnosis of JD by using antigens extracted from the surface of M.  
      avium subsp. paratuberculosis. Nine different chemicals and various  
      intervals of agitation by vortex were evaluated for their ability to  
      extract the surface antigens. Various quantities of surface antigens per  
      well in a 96-well microtiter plate were also tested. The greatest  
      differences in distinguishing between JD-positive and JD-negative serum  
      samples by ethanol vortex enzyme-linked immunosorbent assay (EVELISA) were  
      obtained with surface antigens dislodged from 50 mu g/well of bacilli  
      treated with 80% ethanol followed by a 30-second interval of agitation by  
      vortex. The diagnostic specificity and sensitivity of the EVELISA were  
      97.4% and 100%, respectively. EVELISA plates that had been vacuum-sealed  
      and then tested 7 weeks later (the longest interval tested) had diagnostic  
      specificity and sensitivity rates of 96.9 and 100%, respectively. In a  
      comparative study involving serum samples from 64 fecal culture-positive  
      cattle, the EVELISA identified 96.6% of the low-level fecal shedders and  
      100% of the midlevel and high-level shedders, whereas the Biocor ELISA  
      detected 13.7% of the low-level shedders, 25% of the mid-level shedders,  
      and 96.2% of the high-level shedders. Thus, the EVELISA was substantially  
      superior to the Biocor ELISA, especially in detecting low-level and  
      midlevel shedders. The EVELISA may form the basis for a highly sensitive  
      and subspecies-specific test for the diagnosis of JD.  
 AU   Eda, Shigetoshi; Bannantine, John P.; Waters, W. R.;   \*\*\*Mori,\*\*\*  
 \*\*\*       Yasuyuki\*\*\*   ; Whitlock, Robert H.; Scott, M. Cathy; Speer, C. A.  
 [Reprint  
   Author]  
 AB   Johne's disease (JD), or paratuberculosis, caused by   \*\*\*Mycobacterium\*\*\*  
      avium subsp. paratuberculosis, is one of the most widespread and  
      economically important diseases of livestock and wild ruminants worldwide.  
      Control. . .  
 ORGN . . .  
 Organism Name  
      bovine (common): host, cattle, female  
 Taxa Notes  
      Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates,  
      Nonhuman Mammals, Vertebrates  
 ORGN Classifier

\*\*\*Mycobacteriaceae\*\*\* 08881

Super Taxa

\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;

Eubacteria; Bacteria; Microorganisms

Organism Name

\*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4 ANSWER 5 OF 15 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
DUPLICATE 4
- AN 2006:467815 BIOSIS <<LOGINID::20080325>>
- DN PREV200600465331
- TI A novel enzyme-linked immunosorbent assay for diagnosis of  
\*\*\*Mycobacterium\*\*\* avium subsp paratuberculosis infections (Johne's  
disease) in cattle.
- AU Speer, C. A. [Reprint Author]; Scott, M. Cathy; Bannantine, John P.;  
Waters, W. Ray; \*\*\*Mori, Yasuyuki\*\*\* ; Whitlock, Robert H.; Eda,  
Shigetoshi
- CS Univ Tennessee, Dept Forestry Wildlife and Fisheries, Ctr Wildlife Hlth,  
POB 1071, Knoxville, TN 37901 USA  
caspeer@utk.edu
- SO Clinical and Vaccine Immunology, (MAY 2006) Vol. 13, No. 5, pp. 535-540.  
ISSN: 1556-6811.
- DT Article
- LA English
- ED Entered STN: 20 Sep 2006  
Last Updated on STN: 20 Sep 2006
- AB Enzyme-linked immunosorbent assays (ELISAs) for the diagnosis of Johne's  
disease (JD), caused by \*\*\*Mycobacterium\*\*\* avium subsp.  
paratuberculosis, were developed using whole bacilli treated with  
formaldehyde (called WELISA) or surface antigens obtained by treatment of  
H. avium subsp. paratuberculosis bacilli with formaldehyde and then brief  
sonication (called SELISA). ELISA plates were coated with either whole  
bacilli or sonicated antigens and tested for reactivity against serum  
obtained from JD-positive and JD-negative cattle or from calves  
experimentally inoculated with M. avium subsp. paratuberculosis,  
\*\*\*Mycobacterium\*\*\* avium subsp. avium, or \*\*\*Mycobacterium\*\*\*  
bovis. Because the initial results obtained from the WELISA and SELISA  
were similar, most of the subsequent experiments reported herein were  
performed using the SELISA method. To optimize the SELISA test, various  
concentrations (3.7 to 37%) of formaldehyde and intervals of sonication (2  
to 300 s) were tested. With an increase in formaldehyde concentration and  
a decreased interval of sonication, there was a concomitant decrease in  
nonspecific binding by the SELISA. SELISAs prepared by treating M. avium  
subsp. paratuberculosis with 37% formaldehyde and then a 2-s burst of  
sonication produced the greatest difference (7X) between M. avium subsp.  
paratuberculosis-negative and M. avium subsp. paratuberculosis-positive  
serum samples. The diagnostic sensitivity and specificity for JD by the  
SELISA were greater than 95%. The SELISA showed subspecies-specific  
detection of M. avium subsp. paratuberculosis infections in calves  
experimentally inoculated with M. avium subsp. paratuberculosis or other  
\*\*\*mycobacteria\*\*\*. Based on diagnostic sensitivity and specificity,  
the SELISA appears superior to the commercial ELISAs routinely used for  
the diagnosis of JD.
- TI A novel enzyme-linked immunosorbent assay for diagnosis of  
\*\*\*Mycobacterium\*\*\* avium subsp paratuberculosis infections (Johne's

disease) in cattle.

AU Speer, C. A. [Reprint Author]; Scott, M. Cathy; Bannantine, John P.;  
Waters, W. Ray; \*\*\*Mori, Yasuyuki\*\*\* ; Whitlock, Robert H.; Eda,  
Shigetoshi

AB Enzyme-linked immunosorbent assays (ELISAs) for the diagnosis of Johne's  
disease (JD), caused by \*\*\*Mycobacterium\*\*\* avium subsp.  
paratuberculosis, were developed using whole bacilli treated with  
formaldehyde (called WELISA) or surface antigens obtained by treatment of.  
. . for reactivity against serum obtained from JD-positive and  
JD-negative cattle or from calves experimentally inoculated with M. avium  
subsp. paratuberculosis, \*\*\*Mycobacterium\*\*\* avium subsp. avium, or  
\*\*\*Mycobacterium\*\*\* bovis. Because the initial results obtained from  
the WELISA and SELISA were similar, most of the subsequent experiments  
reported herein. . . showed subspecies-specific detection of M. avium  
subsp. paratuberculosis infections in calves experimentally inoculated  
with M. avium subsp. paratuberculosis or other \*\*\*mycobacteria\*\*\* .  
Based on diagnostic sensitivity and specificity, the SELISA appears  
superior to the commercial ELISAs routinely used for the diagnosis of. .  
.

IT Major Concepts  
Infection; Methods and Techniques

IT Diseases  
\*\*\*Mycobacterium\*\*\* avium tuberculosis infection: bacterial  
disease, infectious disease, diagnosis, Johne's disease

IT Chemicals & Biochemicals  
formaldehyde; surface antigen

ORGN . . .  
Chordata; Animalia  
Organism Name  
cattle (common): host  
Taxa Notes  
Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates,  
Nonhuman Mammals, Vertebrates

ORGN Classifier  
\*\*\*Mycobacteriaceae\*\*\* 08881  
Super Taxa  
\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
Eubacteria; Bacteria; Microorganisms  
Organism Name  
\*\*\*Mycobacterium\*\*\* bovis (species)  
\*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen  
\*\*\*Mycobacterium\*\*\* avium avium (subspecies)  
Taxa Notes  
Bacteria, Eubacteria, Microorganisms

L4 ANSWER 6 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2005:283672 CAPLUS <<LOGINID::20080325>>

DN 142:334896

TI Method for diagnosing johne's disease

IN Momotani, Eiichi; \*\*\*Mori, Yasuyuki\*\*\* ; Hikono, Hirokazu; Buza, Joram  
Josephat

PA Incorporated Administrative Agency National Agriculture and Bio-Oriented  
Research Organization, Japan

SO PCT Int. Appl., 38 pp.  
CODEN: PIXXD2

DT Patent

LA Japanese



FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	----	-----	-----	-----
PI	WO 2005029079	A1	20050331	WO 2003-JP11845	20030917
	W: AU, JP, US				
	RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR				
	AU 2003272880	A1	20050411	AU 2003-272880	20030917
	US 2008038758	A1	20080214	US 2007-572514	20070426
PRAI	WO 2003-JP11845	A	20030917		

AB A method for diagnosing johne's disease is provided, with which an animal infected with \*\*\*Mycobacterium\*\*\* paratuberculosis (Johne's) can be diagnosed at a high sensitivity in the inapparent infection stage before the specific antibody level begins to increase, and a large no. of specimens can be treated. The method is characterized in that it comprises collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* paratuberculosis antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood. The method is also characterized in that the IFN.gamma. yield in blood is measured by the IFN.gamma. ELISA method. Also provided is a method for diagnosing \*\*\*mycobacteriosis\*\*\*, which is characterized by comprising collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood.

RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

IN Momotani, Eiichi; \*\*\*Mori, Yasuyuki\*\*\* ; Hikono, Hirokazu; Buza, Joram Josephat

AB A method for diagnosing johne's disease is provided, with which an animal infected with \*\*\*Mycobacterium\*\*\* paratuberculosis (Johne's) can be diagnosed at a high sensitivity in the inapparent infection stage before the specific antibody level begins. . . is characterized in that it comprises collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* paratuberculosis antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood. The. . . that the IFN.gamma. yield in blood is measured by the IFN.gamma. ELISA method. Also provided is a method for diagnosing \*\*\*mycobacteriosis\*\*\*, which is characterized by comprising collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood.

IT Animals

Blood analysis

Diagnosis

\*\*\*Mycobacterium\*\*\* avium paratuberculosis

(method for diagnosing johne's disease by measuring blood IFN.gamma. by ELISA)

L4 ANSWER 7 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2005:315731 CAPLUS <<LOGINID::20080325>>

DN 142:390942

TI Protein and DNA sequence of \*\*\*Mycobacterium\*\*\* johnei antigens able to induce interferon and uses in diagnosis

IN \*\*\*Mori, Yasuyuki\*\*\* ; Nagata, Reiko; Yoshihara, Kazuhiro; Sota, Yoshihiro; Yokomizo, Yuichi

PA National Institute of Agro-Environmental Sciences, Japan  
 SO Jpn. Kokai Tokkyo Koho, 12 pp.  
 CODEN: JKXXAF  
 DT Patent  
 LA Japanese  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	----	-----	-----	-----
PI	JP 2005095101	A	20050414	JP 2003-334977	20030926
	JP 3864230	B2	20061227		
PRAI	JP 2003-334977		20030926		
AB	The sequences of antigens able to induce interferon .gamma. are isolated from cow PBMC (peripheral blood mononuclear cell) infected with ***Mycobacterium*** johnei. The induction of interferon .gamma. by ***Mycobacterium*** johnei is useful in diagnosis of infection of ***Mycobacterium*** johnei by detection of interferon .gamma. in the supernatant of infected cells.				
TI	Protein and DNA sequence of ***Mycobacterium*** johnei antigens able to induce interferon and uses in diagnosis				
IN	***Mori, Yasuyuki*** ; Nagata, Reiko; Yoshihara, Kazuhiro; Sota, Yoshihiro; Yokomizo, Yuichi				
AB	The sequences of antigens able to induce interferon .gamma. are isolated from cow PBMC (peripheral blood mononuclear cell) infected with ***Mycobacterium*** johnei. The induction of interferon .gamma. by ***Mycobacterium*** johnei is useful in diagnosis of infection of ***Mycobacterium*** johnei by detection of interferon .gamma. in the supernatant of infected cells.				
ST	***Mycobacterium*** antigen sequence interferon gamma induction				
IT	Mononuclear cell (leukocyte) (PBMC; protein and DNA sequence of ***Mycobacterium*** johnei antigens able to induce interferon and uses in diagnosis)				
IT	Animal cell (diagnostic sample from; protein and DNA sequence of ***Mycobacterium*** johnei antigens able to induce interferon and uses in diagnosis)				
IT	Gene, microbial RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study) (encoding antigen; protein and DNA sequence of ***Mycobacterium*** johnei antigens able to induce interferon and uses in diagnosis)				
IT	Diagnosis (mol.; protein and DNA sequence of ***Mycobacterium*** johnei antigens able to induce interferon and uses in diagnosis)				
IT	Infection (of ***Mycobacterium*** johnei, diagnosis of; protein and DNA sequence of ***Mycobacterium*** johnei antigens able to induce interferon and uses in diagnosis)				
IT	Bos taurus DNA sequences ***Mycobacterium*** avium paratuberculosis Protein sequences (protein and DNA sequence of ***Mycobacterium*** johnei antigens able to induce interferon and uses in diagnosis)				
IT	Antigens RL: BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); BIOL (Biological study); USES (Uses) (protein and DNA sequence of ***Mycobacterium*** johnei antigens				

able to induce interferon and uses in diagnosis)

IT Interferons  
 RL: BSU (Biological study, unclassified); DGN (Diagnostic use); BIOL (Biological study); USES (Uses)  
 (.gamma., detection of, in cell supernatant; protein and DNA sequence of \*\*\*Mycobacterium\*\*\* johnei antigens able to induce interferon and uses in diagnosis)

IT 849989-44-2 849989-47-5  
 RL: BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); BIOL (Biological study); USES (Uses)  
 (amino acid sequence; protein and DNA sequence of \*\*\*Mycobacterium\*\*\* johnei antigens able to induce interferon and uses in diagnosis)

IT 849989-45-3 849989-46-4  
 RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (nucleotide sequence; protein and DNA sequence of \*\*\*Mycobacterium\*\*\* johnei antigens able to induce interferon and uses in diagnosis)

L4 ANSWER 8 OF 15 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 5

AN 2005:337763 BIOSIS <<LOGINID::20080325>>

DN PREV200510123867

TI Expression cloning of gamma interferon-inducing antigens of \*\*\*Mycobacterium\*\*\* avium subsp paratuberculosis.

AU Nagata, Reiko [Reprint Author]; Muneta, Yoshihiro; Yoshihara, Kazuhiro; Yokomizo, Yuichi; \*\*\*Mori, Yasuyuki\*\*\*

CS Natl Inst Anim Hlth, Immune Syst Sect, Dept Immunol, 3-1-5 Kannondai, Tsukuba, Ibaraki 3050856, Japan  
 kikuma@affrc.go.jp

SO Infection and Immunity, (JUN 2005) Vol. 73, No. 6, pp. 3778-3782.  
 CODEN: INFIBR. ISSN: 0019-9567.

DT Article

LA English

OS GenBank-AX094821; EMBL-AX094821; DDJB-AX094821; GenBank-U18263; EMBL-U18263; DDJB-U18263

ED Entered STN: 31 Aug 2005  
 Last Updated on STN: 31 Aug 2005

AB Three recombinant proteins, Map10, Map39, and Map41, produced based on nucleotide sequences obtained from the screening of \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis genomic library expressed in Escherichia coli significantly elicited gamma interferon production in peripheral blood mononuclear cells from infected cattle. Two of these proteins were members of the PPE protein family.

TI Expression cloning of gamma interferon-inducing antigens of \*\*\*Mycobacterium\*\*\* avium subsp paratuberculosis.

AU Nagata, Reiko [Reprint Author]; Muneta, Yoshihiro; Yoshihara, Kazuhiro; Yokomizo, Yuichi; \*\*\*Mori, Yasuyuki\*\*\*

AB Three recombinant proteins, Map10, Map39, and Map41, produced based on nucleotide sequences obtained from the screening of \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis genomic library expressed in Escherichia coli significantly elicited gamma interferon production in peripheral blood mononuclear cells from. . .

ORGN . . .  
 Anaerobic Gram-Negative Rods; Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 Escherichia coli (species): expression system  
 Taxa Notes

Bacteria, Eubacteria, Microorganisms  
 ORGN Classifier  
     \*\*\*Mycobacteriaceae\*\*\*      08881  
 Super Taxa  
     \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
     Eubacteria; Bacteria; Microorganisms  
 Organism Name  
     \*\*\*Mycobacterium\*\*\*      avium paratuberculosis (subspecies): pathogen  
 Taxa Notes  
     Bacteria, Eubacteria, Microorganisms

L4 ANSWER 9 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN  
 AN 2004:175700 CAPLUS <<LOGINID::20080325>>  
 DN 140:230513  
 TI Primer sets for detection of \*\*\*Mycobacterium\*\*\* avium and their uses  
     for diagnosis of Johne's disease  
 IN Kageyama, Soichi; Sawai, Takeshi; Hinosawa, Masaki; Onoe, Sadao; Watanabe,  
     Keiko; \*\*\*Mori, Yasuyuki\*\*\* ; Yoshihara, Kazuhiro; Muneta, Yoshihiro;  
     Yokomizo, Yuichi  
 PA Hokkaido Prefecture, Japan; Eiken Chemical Co., Ltd.; Nogyo Gijutsu Kenkyu  
     Kiko  
 SO Jpn. Kokai Tokkyo Koho, 34 pp.  
     CODEN: JKXXAF  
 DT Patent  
 LA Japanese  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 2004065244	A	20040304	JP 2003-159573	20030604
PRAI	JP 2002-168696	A	20020610		

AB This invention provides primer sets for detection of \*\*\*Mycobacterium\*\*\*  
 avium Paratuberculosis. The primers were used for amplification of  
     \*\*\*Mycobacterium\*\*\* insertion sequence IS900. The method of detection  
 of \*\*\*Mycobacterium\*\*\* can be used for diagnosis of Johne's disease.  
 TI Primer sets for detection of \*\*\*Mycobacterium\*\*\* avium and their uses  
     for diagnosis of Johne's disease  
 IN Kageyama, Soichi; Sawai, Takeshi; Hinosawa, Masaki; Onoe, Sadao; Watanabe,  
     Keiko; \*\*\*Mori, Yasuyuki\*\*\* ; Yoshihara, Kazuhiro; Muneta, Yoshihiro;  
     Yokomizo, Yuichi  
 AB This invention provides primer sets for detection of \*\*\*Mycobacterium\*\*\*  
 avium Paratuberculosis. The primers were used for amplification of  
     \*\*\*Mycobacterium\*\*\* insertion sequence IS900. The method of detection  
 of \*\*\*Mycobacterium\*\*\* can be used for diagnosis of Johne's disease.  
 ST primer set detection \*\*\*Mycobacterium\*\*\* diagnosis Johne disease  
 IT Insertion sequence  
     RL: ANT (Analyte); BSU (Biological study, unclassified); ANST (Analytical  
     study); BIOL (Biological study)  
     (IS900, amplification of; primer sets for detection of  
     \*\*\*Mycobacterium\*\*\* avium and their uses for diagnosis of Johne's  
     disease)  
 IT Genetic methods  
     (LAMP, for detection of \*\*\*Mycobacterium\*\*\* ; primer sets for  
     detection of \*\*\*Mycobacterium\*\*\* avium and their uses for diagnosis  
     of Johne's disease)  
 IT \*\*\*Mycobacterium\*\*\* avium paratuberculosis  
     (detection of; primer sets for detection of \*\*\*Mycobacterium\*\*\*  
     avium and their uses for diagnosis of Johne's disease)

IT Genetic methods  
(for detection of \*\*\*Mycobacterium\*\*\* ; primer sets for detection of  
\*\*\*Mycobacterium\*\*\* avium and their uses for diagnosis of Johne's  
disease)

IT Primers (nucleic acid)  
RL: BUU (Biological use, unclassified); DGN (Diagnostic use); PRP  
(Properties); BIOL (Biological study); USES (Uses)  
(for detection of \*\*\*Mycobacterium\*\*\* ; primer sets for detection of  
\*\*\*Mycobacterium\*\*\* avium and their uses for diagnosis of Johne's  
disease)

IT Diagnosis  
(mol., of johne's diseases; primer sets for detection of  
\*\*\*Mycobacterium\*\*\* avium and their uses for diagnosis of Johne's  
disease)

IT Infection  
(paratuberculosis, diagnosis of; primer sets for detection of  
\*\*\*Mycobacterium\*\*\* avium and their uses for diagnosis of Johne's  
disease)

IT 668513-71-1 668513-72-2 668513-73-3 668513-74-4 668513-75-5  
668513-76-6 668513-77-7 668513-78-8 668513-79-9 668513-80-2  
668513-81-3 668513-82-4 668513-83-5 668513-84-6 668513-85-7  
668513-86-8 668513-87-9 668513-88-0 668513-89-1 668513-90-4  
668513-91-5 668513-92-6  
RL: BUU (Biological use, unclassified); DGN (Diagnostic use); PRP  
(Properties); BIOL (Biological study); USES (Uses)  
(nucleotide sequence; primer sets for detection of  
\*\*\*Mycobacterium\*\*\* avium and their uses for diagnosis of Johne's  
disease)

IT 668757-91-3 668757-92-4 668757-93-5 668757-94-6 668757-95-7  
668757-96-8 668757-97-9 668757-98-0 668757-99-1 668758-00-7  
668758-01-8 668758-02-9  
RL: PRP (Properties)  
(unclaimed nucleotide sequence; primer sets for detection of  
\*\*\*Mycobacterium\*\*\* avium and their uses for diagnosis of Johne's  
disease)

L4 ANSWER 10 OF 15 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
STN DUPLICATE 6

AN 2004:438665 BIOSIS <<LOGINID::20080325>>

DN PREV200400437489

TI Neutralization of interleukin-10 significantly enhances gamma interferon  
expression in peripheral blood by stimulation with Johnin purified protein  
derivative and by infection with \*\*\*Mycobacterium\*\*\* avium subsp.  
paratuberculosis in experimentally infected cattle with paratuberculosis.

AU Buza, Jorarn J.; Hikono, Hirokazu; \*\*\*Mori, Yasuyuki\*\*\* ; Nagata,  
Reiko; Hirayama, Sachiyo; Bari, Abusaleh M.; Aodon-geril; Shu, Yujing;  
Tsuji, Noriko M.; Momotani, Eiichi [Reprint Author]

CS ParaTB and Inflammatory Bowel Dis Res Team, NIAH, 3-1-5 Kannondai,  
Tsukuba, Ibaraki, 3050856, Japan  
momotani@affrc.go.jp

SO Infection and Immunity, (April 2004) Vol. 72, No. 4, pp. 2425-2428. print.  
ISSN: 0019-9567 (ISSN print).

DT Article

LA English

ED Entered STN: 17 Nov 2004  
Last Updated on STN: 17 Nov 2004

AB Monoclonal antibody neutralization of interleukin-10 (IL-10) increased

Johnin purified protein derivative-induced whole-blood gamma interferon (IFN-gamma) secretion 23-fold and also increased IFN-gamma secretion ninefold following in vitro \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection of peripheral blood mononuclear cells. These results demonstrate the suppressive effect of IL-10 on immune responses to M. avium subsp. paratuberculosis infection in cattle.

TI. . . interleukin-10 significantly enhances gamma interferon expression in peripheral blood by stimulation with Johnin purified protein derivative and by infection with \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in experimentally infected cattle with paratuberculosis.

AU Buza, Jorarn J.; Hikono, Hirokazu; \*\*\*Mori, Yasuyuki\*\*\* ; Nagata, Reiko; Hirayama, Sachiyo; Bari, Abusaleh M.; Aodon-geril; Shu, Yujing; Tsuji, Noriko M.; Momotani, Eiichi [Reprint Author]

AB. . . increased Johnin purified protein derivative-induced whole-blood gamma interferon (IFN-gamma) secretion 23-fold and also increased IFN-gamma secretion ninefold following in vitro \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection of peripheral blood mononuclear cells. These results demonstrate the suppressive effect of IL-10 on immune responses. . .

ORGN . . .  
 Animalia  
 Organism Name  
 cattle (common): immune responses  
 Taxa Notes  
 Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Vertebrates

ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies)  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms

L4 ANSWER 11 OF 15 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN

AN 2005:45686 BIOSIS <<LOGINID::20080325>>

DN PREV200500044914

TI Generation of multinucleated giant cells in vitro from bovine monocytes and macrophages.

AU Yoshihara, Kazuhiro [Reprint Author]; Nagata, Reiko; Muneta, Yoshihiro; Inumaru, Shigeki; Yokomizo, Yuichi; \*\*\*Mori, Yasuyuki\*\*\*

CS Natl Inst Anim Hlth, 3-1-5 Kannondai, Tsukuba, Ibaraki, 3050856, Japan

SO Journal of Veterinary Medical Science, (September 2004) Vol. 66, No. 9, pp. 1065-1069. print.  
 ISSN: 0916-7250 (ISSN print).

DT Article

LA English

ED Entered STN: 26 Jan 2005  
 Last Updated on STN: 26 Jan 2005

AB The generation of multinucleated giant cells (MGC) from cells of the bovine monocyte-macrophage lineage was investigated. Freshly isolated monocytes were incubated with the conditioned medium (CM) of peripheral blood mononuclear cell cultures treated with Concanavalin A for 1-4 days (CM1 to CM4). Only CM1 generated MGC despite similar concentrations of

IFNgamma in all CMs. Nevertheless, MGC formation from monocytes was enhanced by adding either macrophage colony-stimulating factor (M-CSF) or granulocyte-macrophage colony-stimulating factor (GM-CSF), MGC formations from macrophages were observed only when macrophages were cultured with GM-CSF plus CM. These results indicate that several mechanisms to generate MGC from bovine monocytes-macrophage lineage cells exist, and that GM-CSF is a major mediator of MGC formation in cattle.

AU Yoshihara, Kazuhiro [Reprint Author]; Nagata, Reiko; Muneta, Yoshihiro; Inumaru, Shigeki; Yokomizo, Yuichi; \*\*\*Mori, Yasuyuki\*\*\*

ORGN . . . .  
 Chordata; Animalia  
 Organism Name  
 cattle (common): host  
 Taxa Notes  
 Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates,  
 Nonhuman Mammals, Vertebrates

ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms

L4 ANSWER 12 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN  
 AN 2004:885718 CAPLUS <<LOGINID::20080325>>  
 DN 141:363746  
 TI Development of early-stage diagnostic method for Johne disease by using anti-IL-10 antibody  
 AU Momotani, Eiichi; \*\*\*Mori, Yasuyuki\*\*\*  
 CS Natl. Agric. Bio-oriented Res. Org., Natl. Inst. Animal Health, Tsukuba, 305-0856, Japan  
 SO BRAIN Techno News (2004), 105, 18-24  
 CODEN: BTEEEC; ISSN: 1345-5958  
 PB Nogyo, Seibutsukei Tokutei Sangyo Gijutsu Kenkyu Kiko, Seibutsukei Tokutei Sangyo Gijutsu Kenkyu Shien Senta  
 DT Journal; General Review  
 LA Japanese  
 AB A review on early-stage diagnosis of Johne's disease (paratuberculosis) in cattle by modified interferon .gamma. ELISA assay using IL-10 neutralizing antibody, and its effectiveness.  
 AU Momotani, Eiichi; \*\*\*Mori, Yasuyuki\*\*\*  
 IT Bos taurus  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis  
 (early-stage diagnosis method for Johne's disease using anti-IL-10 antibody)

L4 ANSWER 13 OF 15 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
 DUPLICATE 7  
 AN 2004:64047 BIOSIS <<LOGINID::20080325>>  
 DN PREV200400065534  
 TI \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection causes suppression of RANTES, monocyte chemoattractant protein 1, and tumor necrosis factor alpha expression in peripheral blood of experimentally infected cattle.

AU Buza, Joram J.; \*\*\*Mori, Yasuyuki\*\*\* ; Bari, Abusaleh M.; Hikono, Hirokazu; Aodon-geril; Hirayama, Sachiyo; Shu, Yujing; Momotani, Eiichi [Reprint Author]

CS Paratuberculosis and Inflammatory Bowel Disease Research Team, NIAH, 3-1-5 Kan-nondai, Tsukuba, 305-0856, Japan  
momotani@affrc.go.jp

SO Infection and Immunity, (December 2003) Vol. 71, No. 12, pp. 7223-7227. print.  
ISSN: 0019-9567 (ISSN print).

DT Article

LA English

ED Entered STN: 28 Jan 2004  
Last Updated on STN: 28 Jan 2004

AB Blood from cattle with subclinical \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection was stimulated with M. avium subsp. paratuberculosis antigens, and expression of interleukin-1beta (IL-1beta), tumor necrosis factor alpha (TNF-alpha), RANTES, monocyte chemoattractant protein 1 (MCP-1), and IL-8 was measured. Expression of TNF-alpha, RANTES, and MCP-1 was lower in infected than in uninfected cattle. The reduced response may weaken protective immunity and perpetuate infection.

TI \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection causes suppression of RANTES, monocyte chemoattractant protein 1, and tumor necrosis factor alpha expression in peripheral. . .

AU Buza, Joram J.; \*\*\*Mori, Yasuyuki\*\*\* ; Bari, Abusaleh M.; Hikono, Hirokazu; Aodon-geril; Hirayama, Sachiyo; Shu, Yujing; Momotani, Eiichi [Reprint Author]

AB Blood from cattle with subclinical \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection was stimulated with M. avium subsp. paratuberculosis antigens, and expression of interleukin-1beta (IL-1beta), tumor necrosis factor. . .

ORGN . . .  
Animalia  
Organism Name  
cattle (common): host, breed-Holstein  
Taxa Notes  
Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Vertebrates

ORGN Classifier  
\*\*\*Mycobacteriaceae\*\*\* 08881  
Super Taxa  
\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
Eubacteria; Bacteria; Microorganisms  
Organism Name  
\*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen  
Taxa Notes  
Bacteria, Eubacteria, Microorganisms

L4 ANSWER 14 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2003:399194 CAPLUS <<LOGINID::20080325>>

DN 140:39839

TI Studies on diagnostic methods for bovine paratuberculosis

AU \*\*\*Mori, Yasuyuki\*\*\* ; Kikuma, Reiko; Muneta, Yoshihiro; Yoshihara, Kazuhiro; Hikono, Hirokazu; Momotani, Eiichi

CS Immune System Section, Department of Immunology, National Institute of Animal Health, Tsukuba, 305-0856, Japan

SO Dobutsu Eisei Kenkyusho Kenkyu Hokoku (2003), Volume Date 2002, 109, 33-42  
CODEN: DEKKC9; ISSN: 1347-2542



PB Nogyo Gijutsu Kenkyu Kiko Dobutsu Eisei Kenkyusho  
 DT Journal  
 LA Japanese  
 AB Current diagnostic tests for paratuberculosis principally rest on serol. assay, bacterial culture and the johnin skin test. However, diagnostic tests that are both sensitive and specific for detecting all subclinically affected animals have not yet been found. Therefore, a no. of studies have been conducted in order to find rapid and accurate diagnostic methods for paratuberculosis. As a result, the following have been found. (1) PCR test with internal control DNA is accurate, sensitive and rapid for the detection of \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in fecal samples. (2) In the interferon gamma (IFN-.gamma.) assay using johnin purified protein deriv. (J-PPD), bovine tuberculin PPD and Con A (Con A), IFN-.gamma. responses against J-PPD were the highest in affected animals. On the contrary those of Con A were the highest in healthy animals. Interpretation of the IFN-.gamma. assay by the higher IFN-.gamma. responses against J-PPD than those of Con A is preferable as one of the diagnostic criteria. (3) Monoclonal antibody (711-1-1) which recognizes the lipoarabinomannan antigen of M. avium subsp. paratuberculosis did not react with M. avium subsp. avium, and showed potential usefulness in the serol. tests. (4) A recombinant alkyl hydroperoxide reductase C of M. avium subsp. paratuberculosis has been prepd. and successfully applied to induce IFN-.gamma. from peripheral blood mononuclear cells of animals infected with M. avium subsp. paratuberculosis. (5) In the course of study on the role of cytokines, monocyte chemoattractant protein-1 seems to be involved in the pathogenesis of paratuberculosis.

AU \*\*\*Mori, Yasuyuki\*\*\* ; Kikuma, Reiko; Muneta, Yoshihiro; Yoshihara, Kazuhiro; Hikono, Hirokazu; Momotani, Eiichi  
 AB . . . following have been found. (1) PCR test with internal control DNA is accurate, sensitive and rapid for the detection of  
 \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in fecal samples.

(2) In the interferon gamma (IFN-.gamma.) assay using johnin purified protein deriv. (J-PPD), bovine. . .

ST cattle \*\*\*Mycobacterium\*\*\* paratuberculosis infection surface antigen IFN induction test; alkyl hydroperoxide reductase antigen cattle IFN induction \*\*\*Mycobacterium\*\*\* ; reverse transcription PCR monocyte chemoattractant protein mRNA assay

IT Bos taurus  
 Infection  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis  
 (studies on diagnostic methods for bovine paratuberculosis)

L4 ANSWER 15 OF 15 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
 AN 2003:329566 BIOSIS <<LOGINID::20080325>>  
 DN PREV200300329566  
 TI Studies on the diagnostic methods for bovine paratuberculosis.  
 AU \*\*\*Mori, Yasuyuki\*\*\* [Reprint Author]; Kikuma, Reiko; Muneta, Yoshihiro; Yoshihara, Kazuhiro; Hikono, Hirokazu; Momotani, Eiichi  
 CS Immune System Section, Department of Immunology, National Institute of Animal Health, 3-1-5 Kannondai, Tsukuba, Ibaraki, 305-0856, Japan  
 yamori@affrc.go.jp  
 SO Bulletin of the National Institute of Animal Health, (2002) No. 109, pp. 33-42. print.  
 ISSN: 1347-2542 (ISSN print).

DT Article  
 LA Japanese  
 ED Entered STN: 16 Jul 2003  
 Last Updated on STN: 16 Jul 2003  
 AB Current diagnostic tests for paratuberculosis principally rest on serological assay, bacterial culture and the johnin skin test. However, diagnostic tests that are both sensitive and specific for detecting all subclinically affected animals have not yet been found. Therefore, a number of studies have been conducted in order to find rapid and accurate diagnostic methods for paratuberculosis. As a result, the following have been found; 1) PCR test with internal control DNA is accurate, sensitive and rapid for the detection of \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in faecal samples. 2) In the interferon gamma (IFN-gamma) assay using johnin purified protein derivative (J-PPD), bovine tuberculin PPD and concanavalin A (Con A), IFN-gamma responses against J-PPD were the highest in affected animals. On the contrary those of Con A were the highest in healthy animals. Interpretation of the IFN-gamma assay by the higher IFN-gamma responses against J-PPD than those of Con A is preferable as one of the diagnostic criteria. 3) Monoclonal antibody (711-1-1) which recognizes the lipoarabinomannan antigen of M. avium subsp. paratuberculosis did not react with M. avium subsp. avium, and showed potential usefulness in the serological tests. 4) A recombinant alkyl hydroperoxide reductase C of M. avium subsp. paratuberculosis has been prepared and successfully applied to induce IFN-gamma from peripheral blood mononuclear cells of animals infected with M. avium subsp. paratuberculosis. 5) In the course of study on the role of cytokines, monocyte chemoattractant protein-1 seems to be involved in the pathogenesis of paratuberculosis.  
 AU \*\*\*Mori, Yasuyuki\*\*\* [Reprint Author]; Kikuma, Reiko; Muneta, Yoshihiro; Yoshihara, Kazuhiro; Hikono, Hirokazu; Momotani, Eiichi  
 AB. . . following have been found; 1) PCR test with internal control DNA is accurate, sensitive and rapid for the detection of \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in faecal samples. 2) In the interferon gamma (IFN-gamma) assay using johnin purified protein derivative (J-PPD), bovine. . .  
 ORGN . . .  
 Chordata; Animalia  
 Organism Name  
 bovine (common): host  
 Taxa Notes  
 Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Vertebrates  
 ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen, strain-ATCC 19698, strain-Kag-1  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms  
 => e hikono hirokazu/au  
 E1 1 HIKONO ATSUSHI/AU  
 E2 26 HIKONO H/AU

E3 39 --> HIKONO HIROKAZU/AU  
 E4 2 HIKONO KOICHI/AU  
 E5 1 HIKONO M/AU  
 E6 1 HIKONO MASAHARU/AU  
 E7 1 HIKONO MASAJI/AU  
 E8 5 HIKONO TAKIO/AU  
 E9 1 HIKONOV V A/AU  
 E10 21 HIKOSAKA A/AU  
 E11 7 HIKOSAKA AIZO/AU  
 E12 14 HIKOSAKA AKIHIIDE/AU

=> s e2-e3 and mycobact?

L5 13 ("HIKONO H"/AU OR "HIKONO HIROKAZU"/AU) AND MYCOBACT?

=> dup rem l5

PROCESSING COMPLETED FOR L5

L6 5 DUP REM L5 (8 DUPLICATES REMOVED)

=> d bib ab kwic

L6 ANSWER 1 OF 5 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2005:283672 CAPLUS <<LOGINID::20080325>>

DN 142:334896

TI Method for diagnosing johne's disease

IN Momotani, Eiichi; Mori, Yasuyuki; \*\*\*Hikono, Hirokazu\*\*\* ; Buza, Joram Josephat

PA Incorporated Administrative Agency National Agriculture and Bio-Oriented Research Organization, Japan

SO PCT Int. Appl., 38 pp.

CODEN: PIXXD2

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2005029079	A1	20050331	WO 2003-JP11845	20030917
	W: AU, JP, US				
	RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR				
	AU 2003272880	A1	20050411	AU 2003-272880	20030917
	US 2008038758	A1	20080214	US 2007-572514	20070426
PRAI	WO 2003-JP11845	A	20030917		

AB A method for diagnosing johne's disease is provided, with which an animal infected with \*\*\*Mycobacterium\*\*\* paratuberculosis (Johne's) can be diagnosed at a high sensitivity in the inapparent infection stage before the specific antibody level begins to increase, and a large no. of specimens can be treated. The method is characterized in that it comprises collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* paratuberculosis antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood. The method is also characterized in that the IFN.gamma. yield in blood is measured by the IFN.gamma. ELISA method. Also provided is a method for diagnosing \*\*\*mycobacteriosis\*\*\*, which is characterized by comprising collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood.

RE.CNT 3        THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD  
              ALL CITATIONS AVAILABLE IN THE RE FORMAT

IN    Momotani, Eiichi; Mori, Yasuyuki;   \*\*\*Hikono, Hirokazu\*\*\*   ; Buza, Joram  
      Josephat

AB    A method for diagnosing johne's disease is provided, with which an animal  
      infected with   \*\*\*Mycobacterium\*\*\*   paratuberculosis (Johne's) can be  
      diagnosed at a high sensitivity in the inapparent infection stage before  
      the specific antibody level begins. . . is characterized in that it  
      comprises collecting a blood sample of a subject animal, adding an  
      anti-IL-10 antibody and a   \*\*\*Mycobacterium\*\*\*   paratuberculosis antigen  
      to the collected blood followed by culturing, and then, measuring the  
      IFN.gamma. yield in the cultured blood. The. . . that the IFN.gamma.  
      yield in blood is measured by the IFN.gamma. ELISA method. Also provided  
      is a method for diagnosing   \*\*\*mycobacteriosis\*\*\*   , which is  
      characterized by comprising collecting a blood sample of a subject animal,  
      adding an anti-IL-10 antibody and a   \*\*\*Mycobacterium\*\*\*   antigen to the  
      collected blood followed by culturing, and then, measuring the IFN.gamma.  
      yield in the cultured blood.

IT    Animals

      Blood analysis

      Diagnosis

          \*\*\*Mycobacterium\*\*\*   avium paratuberculosis

          (method for diagnosing johne's disease by measuring blood IFN.gamma. by  
          ELISA)

=> d bib ab kwic 2-

YOU HAVE REQUESTED DATA FROM 4 ANSWERS - CONTINUE? Y/(N):y

L6    ANSWER 2 OF 5   BIOSIS   COPYRIGHT (c) 2008 The Thomson Corporation   on STN  
      DUPLICATE 1

AN    2004:438665   BIOSIS <<LOGINID::20080325>>

DN    PREV200400437489

TI    Neutralization of interleukin-10 significantly enhances gamma interferon  
      expression in peripheral blood by stimulation with Johnin purified protein  
      derivative and by infection with   \*\*\*Mycobacterium\*\*\*   avium subsp.  
      paratuberculosis in experimentally infected cattle with paratuberculosis.

AU    Buza, Jorarn J.;   \*\*\*Hikono, Hirokazu\*\*\*   ; Mori, Yasuyuki; Nagata,  
      Reiko; Hirayama, Sachiyo; Bari, Abusaleh M.; Aodon-geril; Shu, Yujing;  
      Tsuji, Noriko M.; Momotani, Eiichi [Reprint Author]

CS    ParaTB and Inflammatory Bowel Dis Res Team, NIAH, 3-1-5 Kannondai,  
      Tsukuba, Ibaraki, 3050856, Japan  
      momotani@affrc.go.jp

SO    Infection and Immunity, (April 2004) Vol. 72, No. 4, pp. 2425-2428. print.  
      ISSN: 0019-9567 (ISSN print).

DT    Article

LA    English

ED    Entered STN: 17 Nov 2004

      Last Updated on STN: 17 Nov 2004

AB    Monoclonal antibody neutralization of interleukin-10 (IL-10) increased  
      Johnin purified protein derivative-induced whole-blood gamma interferon  
      (IFN-gamma) secretion 23-fold and also increased IFN-gamma secretion  
      ninefold following in vitro   \*\*\*Mycobacterium\*\*\*   avium subsp.  
      paratuberculosis infection of peripheral blood mononuclear cells. These  
      results demonstrate the suppressive effect of IL-10 on immune responses to  
      M. avium subsp. paratuberculosis infection in cattle.

TI.   . . interleukin-10 significantly enhances gamma interferon expression in

peripheral blood by stimulation with Johnin purified protein derivative and by infection with \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in experimentally infected cattle with paratuberculosis.

AU Buza, Jorarn J.; \*\*\*Hikono, Hirokazu\*\*\* ; Mori, Yasuyuki; Nagata, Reiko; Hirayama, Sachiyo; Bari, Abusaleh M.; Aodon-geril; Shu, Yujing; Tsuji, Noriko M.; Momotani, Eiichi [Reprint Author]

AB. . . increased Johnin purified protein derivative-induced whole-blood gamma interferon (IFN-gamma) secretion 23-fold and also increased IFN-gamma secretion ninefold following in vitro \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection of peripheral blood mononuclear cells. These results demonstrate the suppressive effect of IL-10 on immune responses. . .

ORGN . . .  
 Animalia  
 Organism Name  
 cattle (common): immune responses  
 Taxa Notes  
 Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Vertebrates

ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies)  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms

L6 ANSWER 3 OF 5 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 2

AN 2004:64047 BIOSIS <<LOGINID::20080325>>

DN PREV200400065534

TI \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection causes suppression of RANTES, monocyte chemoattractant protein 1, and tumor necrosis factor alpha expression in peripheral blood of experimentally infected cattle.

AU Buza, Joram J.; Mori, Yasuyuki; Bari, Abusaleh M.; \*\*\*Hikono,\*\*\*  
 \*\*\* Hirokazu\*\*\* ; Aodon-geril; Hirayama, Sachiyo; Shu, Yujing; Momotani, Eiichi [Reprint Author]

CS Paratuberculosis and Inflammatory Bowel Disease Research Team, NIAH, 3-1-5 Kan-nondai, Tsukuba, 305-0856, Japan  
 momotani@affrc.go.jp

SO Infection and Immunity, (December 2003) Vol. 71, No. 12, pp. 7223-7227. print.  
 ISSN: 0019-9567 (ISSN print).

DT Article

LA English

ED Entered STN: 28 Jan 2004  
 Last Updated on STN: 28 Jan 2004

AB Blood from cattle with subclinical \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection was stimulated with M. avium subsp. paratuberculosis antigens, and expression of interleukin-1beta (IL-1beta), tumor necrosis factor alpha (TNF-alpha), RANTES, monocyte chemoattractant protein 1 (MCP-1), and IL-8 was measured. Expression of TNF-alpha, RANTES, and MCP-1 was lower in infected than in uninfected cattle. The reduced response may weaken protective immunity and perpetuate infection.

TI \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection causes suppression of RANTES, monocyte chemoattractant protein 1, and tumor necrosis factor alpha expression in peripheral. . . .  
 AU Buza, Joram J.; Mori, Yasuyuki; Bari, Abusaleh M.; \*\*\*Hikono,\*\*\*  
 \*\*\* Hirokazu\*\*\* ; Aodon-geril; Hirayama, Sachiyo; Shu, Yujing; Momotani, Eiichi [Reprint Author]  
 AB Blood from cattle with subclinical \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection was stimulated with M. avium subsp. paratuberculosis antigens, and expression of interleukin-1beta (IL-1beta), tumor necrosis factor. . . .  
 ORGN . . . .  
 Animalia  
 Organism Name  
 cattle (common): host, breed-Holstein  
 Taxa Notes  
 Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Vertebrates  
 ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms  
  
 L6 ANSWER 4 OF 5 CAPLUS COPYRIGHT 2008 ACS on STN  
 AN 2003:399194 CAPLUS <<LOGINID::20080325>>  
 DN 140:39839  
 TI Studies on diagnostic methods for bovine paratuberculosis  
 AU Mori, Yasuyuki; Kikuma, Reiko; Muneta, Yoshihiro; Yoshihara, Kazuhiro;  
 \*\*\*Hikono, Hirokazu\*\*\* ; Momotani, Eiichi  
 CS Immune System Section, Department of Immunology, National Institute of Animal Health, Tsukuba, 305-0856, Japan  
 SO Dobutsu Eisei Kenkyusho Kenkyu Hokoku (2003), Volume Date 2002, 109, 33-42  
 CODEN: DEKKC9; ISSN: 1347-2542  
 PB Nogyo Gijutsu Kenkyu Kiko Dobutsu Eisei Kenkyusho  
 DT Journal  
 LA Japanese  
 AB Current diagnostic tests for paratuberculosis principally rest on serol. assay, bacterial culture and the johnin skin test. However, diagnostic tests that are both sensitive and specific for detecting all subclinically affected animals have not yet been found. Therefore, a no. of studies have been conducted in order to find rapid and accurate diagnostic methods for paratuberculosis. As a result, the following have been found. (1) PCR test with internal control DNA is accurate, sensitive and rapid for the detection of \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis in fecal samples. (2) In the interferon gamma (IFN-.gamma.) assay using johnin purified protein deriv. (J-PPD), bovine tuberculin PPD and Con A (Con A), IFN-.gamma. responses against J-PPD were the highest in affected animals. On the contrary those of Con A were the highest in healthy animals. Interpretation of the IFN-.gamma. assay by the higher IFN-.gamma. responses against J-PPD than those of Con A is preferable as one of the diagnostic criteria. (3) Monoclonal antibody (711-1-1) which recognizes the lipoarabinomannan antigen of M. avium subsp. paratuberculosis did not react with M. avium subsp. avium, and showed

potential usefulness in the serol. tests. (4) A recombinant alkyl hydroperoxide reductase C of *M. avium* subsp. *paratuberculosis* has been prepd. and successfully applied to induce IFN- $\gamma$  from peripheral blood mononuclear cells of animals infected with *M. avium* subsp. *paratuberculosis*. (5) In the course of study on the role of cytokines, monocyte chemoattractant protein-1 seems to be involved in the pathogenesis of *paratuberculosis*.

AU Mori, Yasuyuki; Kikuma, Reiko; Muneta, Yoshihiro; Yoshihara, Kazuhiro;  
 \*\*\*Hikono, Hirokazu\*\*\* ; Momotani, Eiichi

AB . . . following have been found. (1) PCR test with internal control DNA is accurate, sensitive and rapid for the detection of  
 \*\*\*Mycobacterium\*\*\* *avium* subsp. *paratuberculosis* in fecal samples.

(2) In the interferon gamma (IFN- $\gamma$ ) assay using johnin purified protein deriv. (J-PPD), bovine. . .

ST cattle \*\*\*Mycobacterium\*\*\* *paratuberculosis* infection surface antigen IFN induction test; alkyl hydroperoxide reductase antigen cattle IFN induction \*\*\*Mycobacterium\*\*\* ; reverse transcription PCR monocyte chemoattractant protein mRNA assay

IT Bos taurus  
 Infection  
 \*\*\*Mycobacterium\*\*\* *avium paratuberculosis*  
 (studies on diagnostic methods for bovine *paratuberculosis*)

L6 ANSWER 5 OF 5 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 3

AN 2003:329566 BIOSIS <<LOGINID::20080325>>

DN PREV200300329566

TI Studies on the diagnostic methods for bovine *paratuberculosis*.

AU Mori, Yasuyuki [Reprint Author]; Kikuma, Reiko; Muneta, Yoshihiro; Yoshihara, Kazuhiro; \*\*\*Hikono, Hirokazu\*\*\* ; Momotani, Eiichi

CS Immune System Section, Department of Immunology, National Institute of Animal Health, 3-1-5 Kannondai, Tsukuba, Ibaraki, 305-0856, Japan  
 yamori@affrc.go.jp

SO Bulletin of the National Institute of Animal Health, (2002) No. 109, pp. 33-42. print.  
 ISSN: 1347-2542 (ISSN print).

DT Article

LA Japanese

ED Entered STN: 16 Jul 2003  
 Last Updated on STN: 16 Jul 2003

AB Current diagnostic tests for *paratuberculosis* principally rest on serological assay, bacterial culture and the johnin skin test. However, diagnostic tests that are both sensitive and specific for detecting all subclinically affected animals have not yet been found. Therefore, a number of studies have been conducted in order to find rapid and accurate diagnostic methods for *paratuberculosis*. As a result, the following have been found; 1) PCR test with internal control DNA is accurate, sensitive and rapid for the detection of \*\*\*Mycobacterium\*\*\* *avium* subsp. *paratuberculosis* in faecal samples. 2) In the interferon gamma (IFN- $\gamma$ ) assay using johnin purified protein derivative (J-PPD), bovine tuberculin PPD and concanavalin A (Con A), IFN- $\gamma$  responses against J-PPD were the highest in affected animals. On the contrary those of Con A were the highest in healthy animals. Interpretation of the IFN- $\gamma$  assay by the higher IFN- $\gamma$  responses against J-PPD than those of Con A is preferable as one of the diagnostic criteria. 3) Monoclonal antibody (711-1-1) which recognizes the lipoarabinomannan antigen of *M. avium* subsp.

paratuberculosis did not react with *M. avium* subsp. *avium*, and showed potential usefulness in the serological tests. 4) A recombinant alkyl hydroperoxide reductase C of *M. avium* subsp. *paratuberculosis* has been prepared and successfully applied to induce IFN-gamma from peripheral blood mononuclear cells of animals infected with *M. avium* subsp. *paratuberculosis*. 5) In the course of study on the role of cytokines, monocyte chemoattractant protein-1 seems to be involved in the pathogenesis of paratuberculosis.

AU Mori, Yasuyuki [Reprint Author]; Kikuma, Reiko; Muneta, Yoshihiro; Yoshihara, Kazuhiro; \*\*\*Hikono, Hirokazu\*\*\* ; Momotani, Eiichi

AB. . . following have been found; 1) PCR test with internal control DNA is accurate, sensitive and rapid for the detection of \*\*\**Mycobacterium*\*\*\* *avium* subsp. *paratuberculosis* in faecal samples. 2) In the interferon gamma (IFN-gamma) assay using johnin purified protein derivative (J-PPD), bovine. . .

ORGN . . .

Chordata; Animalia

Organism Name

bovine (common): host

Taxa Notes

Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Vertebrates

ORGN Classifier

\*\*\*Mycobacteriaceae\*\*\* 08881

Super Taxa

\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;

Eubacteria; Bacteria; Microorganisms

Organism Name

\*\*\*Mycobacterium\*\*\* *avium paratuberculosis* (subspecies): pathogen, strain-ATCC 19698, strain-Kag-1

Taxa Notes

Bacteria, Eubacteria, Microorganisms

=> e buza joram j/au

E1	11	BUZA J J/AU
E2	3	BUZA JORAM/AU
E3	8	--> BUZA JORAM J/AU
E4	1	BUZA JORAM JOSEPHAT/AU
E5	1	BUZA JORARN J/AU
E6	4	BUZA K/AU
E7	19	BUZA L/AU
E8	1	BUZA L N/AU
E9	1	BUZA L V/AU
E10	7	BUZA LAJOSNE/AU
E11	2	BUZA LASZLO/AU
E12	1	BUZA LEJLA/AU

=> s e1-e5 and mycobact?

L7 11 ("BUZA J J"/AU OR "BUZA JORAM"/AU OR "BUZA JORAM J"/AU OR "BUZA JORAM JOSEPHAT"/AU OR "BUZA JORARN J"/AU) AND MYCOBACT?

=> dup rem l7

PROCESSING COMPLETED FOR L7

L8 3 DUP REM L7 (8 DUPLICATES REMOVED)

=> d bib ab kwic 1-



YOU HAVE REQUESTED DATA FROM 3 ANSWERS - CONTINUE? Y/(N):y

L8 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2005:283672 CAPLUS <<LOGINID::20080325>>

DN 142:334896

TI Method for diagnosing johne's disease

IN Momotani, Eiichi; Mori, Yasuyuki; Hikono, Hirokazu; \*\*\*Buza, Joram\*\*\*

\*\*\* Josephat\*\*\*

PA Incorporated Administrative Agency National Agriculture and Bio-Oriented  
Research Organization, Japan

SO PCT Int. Appl., 38 pp.

CODEN: PIXXD2

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	----	-----	-----	-----
PI	WO 2005029079	A1	20050331	WO 2003-JP11845	20030917
	W: AU, JP, US				
	RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR				
	AU 2003272880	A1	20050411	AU 2003-272880	20030917
	US 2008038758	A1	20080214	US 2007-572514	20070426
PRAI	WO 2003-JP11845	A	20030917		

AB A method for diagnosing johne's disease is provided, with which an animal infected with \*\*\*Mycobacterium\*\*\* paratuberculosis (Johne's) can be diagnosed at a high sensitivity in the inapparent infection stage before the specific antibody level begins to increase, and a large no. of specimens can be treated. The method is characterized in that it comprises collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* paratuberculosis antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood. The method is also characterized in that the IFN.gamma. yield in blood is measured by the IFN.gamma. ELISA method. Also provided is a method for diagnosing \*\*\*mycobacteriosis\*\*\*, which is characterized by comprising collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood.

RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

IN Momotani, Eiichi; Mori, Yasuyuki; Hikono, Hirokazu; \*\*\*Buza, Joram\*\*\*

\*\*\* Josephat\*\*\*

AB A method for diagnosing johne's disease is provided, with which an animal infected with \*\*\*Mycobacterium\*\*\* paratuberculosis (Johne's) can be diagnosed at a high sensitivity in the inapparent infection stage before the specific antibody level begins. . . is characterized in that it comprises collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* paratuberculosis antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood. The. . . that the IFN.gamma. yield in blood is measured by the IFN.gamma. ELISA method. Also provided is a method for diagnosing \*\*\*mycobacteriosis\*\*\*, which is characterized by comprising collecting a blood sample of a subject animal, adding an anti-IL-10 antibody and a \*\*\*Mycobacterium\*\*\* antigen to the collected blood followed by culturing, and then, measuring the IFN.gamma. yield in the cultured blood.

IT Animals  
Blood analysis  
Diagnosis  
\*\*\*Mycobacterium\*\*\* avium paratuberculosis  
(method for diagnosing johne's disease by measuring blood IFN.gamma. by ELISA)

L8 ANSWER 2 OF 3 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
DUPLICATE 1

AN 2004:438665 BIOSIS <<LOGINID::20080325>>  
DN PREV200400437489

TI Neutralization of interleukin-10 significantly enhances gamma interferon  
expression in peripheral blood by stimulation with Johnin purified protein  
derivative and by infection with \*\*\*Mycobacterium\*\*\* avium subsp.  
paratuberculosis in experimentally infected cattle with paratuberculosis.

AU \*\*\*Buza, Jorarn J.\*\*\* ; Hikono, Hirokazu; Mori, Yasuyuki; Nagata,  
Reiko; Hirayama, Sachiyo; Bari, Abusaleh M.; Aodon-geril; Shu, Yujing;  
Tsuji, Noriko M.; Momotani, Eiichi [Reprint Author]

CS ParaTB and Inflammatory Bowel Dis Res Team, NIAH, 3-1-5 Kannondai,  
Tsukuba, Ibaraki, 3050856, Japan  
momotani@affrc.go.jp

SO Infection and Immunity, (April 2004) Vol. 72, No. 4, pp. 2425-2428. print.  
ISSN: 0019-9567 (ISSN print).

DT Article  
LA English  
ED Entered STN: 17 Nov 2004  
Last Updated on STN: 17 Nov 2004

AB Monoclonal antibody neutralization of interleukin-10 (IL-10) increased  
Johnin purified protein derivative-induced whole-blood gamma interferon  
(IFN-gamma) secretion 23-fold and also increased IFN-gamma secretion  
ninefold following in vitro \*\*\*Mycobacterium\*\*\* avium subsp.  
paratuberculosis infection of peripheral blood mononuclear cells. These  
results demonstrate the suppressive effect of IL-10 on immune responses to  
M. avium subsp. paratuberculosis infection in cattle.

TI. . . interleukin-10 significantly enhances gamma interferon expression in  
peripheral blood by stimulation with Johnin purified protein derivative  
and by infection with \*\*\*Mycobacterium\*\*\* avium subsp.  
paratuberculosis in experimentally infected cattle with paratuberculosis.

AU \*\*\*Buza, Jorarn J.\*\*\* ; Hikono, Hirokazu; Mori, Yasuyuki; Nagata,  
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Tsuji, Noriko M.; Momotani, . . .

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IFN-gamma secretion ninefold following in vitro \*\*\*Mycobacterium\*\*\*  
avium subsp. paratuberculosis infection of peripheral blood mononuclear  
cells. These results demonstrate the suppressive effect of IL-10 on  
immune responses. . .

ORGN . . .  
Animalia  
Organism Name  
cattle (common): immune responses  
Taxa Notes  
Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates,  
Nonhuman Mammals, Vertebrates

ORGN Classifier  
\*\*\*Mycobacteriaceae\*\*\* 08881  
Super Taxa

\*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms  
 Organism Name  
 \*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies)  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms

L8 ANSWER 3 OF 3 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
 DUPLICATE 2  
 AN 2004:64047 BIOSIS <<LOGINID::20080325>>  
 DN PREV200400065534  
 TI \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection causes  
 suppression of RANTES, monocyte chemoattractant protein 1, and tumor  
 necrosis factor alpha expression in peripheral blood of experimentally  
 infected cattle.  
 AU \*\*\*Buza, Joram J.\*\*\* ; Mori, Yasuyuki; Bari, Abusaleh M.; Hikono,  
 Hirokazu; Aodon-geril; Hirayama, Sachiyo; Shu, Yujing; Momotani, Eiichi  
 [Reprint Author]  
 CS Paratuberculosis and Inflammatory Bowel Disease Research Team, NIAH, 3-1-5  
 Kan-nondai, Tsukuba, 305-0856, Japan  
 momotani@affrc.go.jp  
 SO Infection and Immunity, (December 2003) Vol. 71, No. 12, pp. 7223-7227.  
 print.  
 ISSN: 0019-9567 (ISSN print).  
 DT Article  
 LA English  
 ED Entered STN: 28 Jan 2004  
 Last Updated on STN: 28 Jan 2004  
 AB Blood from cattle with subclinical \*\*\*Mycobacterium\*\*\* avium subsp.  
 paratuberculosis infection was stimulated with M. avium subsp.  
 paratuberculosis antigens, and expression of interleukin-1beta (IL-1beta),  
 tumor necrosis factor alpha (TNF-alpha), RANTES, monocyte chemoattractant  
 protein 1 (MCP-1), and IL-8 was measured. Expression of TNF-alpha,  
 RANTES, and MCP-1 was lower in infected than in uninfected cattle. The  
 reduced response may weaken protective immunity and perpetuate infection.  
 TI \*\*\*Mycobacterium\*\*\* avium subsp. paratuberculosis infection causes  
 suppression of RANTES, monocyte chemoattractant protein 1, and tumor  
 necrosis factor alpha expression in peripheral. . .  
 AU \*\*\*Buza, Joram J.\*\*\* ; Mori, Yasuyuki; Bari, Abusaleh M.; Hikono,  
 Hirokazu; Aodon-geril; Hirayama, Sachiyo; Shu, Yujing; Momotani, Eiichi  
 [Reprint Author]  
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 paratuberculosis infection was stimulated with M. avium subsp.  
 paratuberculosis antigens, and expression of interleukin-1beta (IL-1beta),  
 tumor necrosis factor. . .  
 ORGN . . .  
 Animalia  
 Organism Name  
 cattle (common): host, breed-Holstein  
 Taxa Notes  
 Animals, Artiodactyls, Chordates, Mammals, Nonhuman Vertebrates,  
 Nonhuman Mammals, Vertebrates  
 ORGN Classifier  
 \*\*\*Mycobacteriaceae\*\*\* 08881  
 Super Taxa  
 \*\*\*Mycobacteria\*\*\* ; Actinomycetes and Related Organisms;  
 Eubacteria; Bacteria; Microorganisms

Organism Name  
\*\*\*Mycobacterium\*\*\* avium paratuberculosis (subspecies): pathogen  
Taxa Notes  
Bacteria, Eubacteria, Microorganisms

=> s paratuberculosis and interleukin? and interferon? and (diagnos? or assay?)  
L9 36 PARATUBERCULOSIS AND INTERLEUKIN? AND INTERFERON? AND (DIAGNOS?  
OR ASSAY?)

=> dup rem l9  
PROCESSING COMPLETED FOR L9  
L10 19 DUP REM L9 (17 DUPLICATES REMOVED)

=> d bib ab kwic 1-  
YOU HAVE REQUESTED DATA FROM 19 ANSWERS - CONTINUE? Y/(N):y

L10 ANSWER 1 OF 19 CAPLUS COPYRIGHT 2008 ACS on STN  
AN 2007:906779 CAPLUS <<LOGINID::20080325>>  
DN 147:275692  
TI Sequences for Mycobacterium leprae-specific antigens, and methods for  
treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in the early  
stages and paucibacillary infections  
IN Ottenhof, Tom Henricus Maria; Geluk, Annemieke; Pereira Sampaio, Elizabeth  
PA Leiden University Medical Center, Neth.  
SO PCT Int. Appl., 70pp.  
CODEN: PIXXD2  
DT Patent  
LA English  
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	----	-----	-----	-----
PI	WO 2007091881	A2	20070816	WO 2006-NL50105	20060428
	WO 2007091881	A3	20071129		
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
	RW:	AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AP, EA, EP, OA			

PRAI EP 2005-103576 A 20050429

AB The current invention discloses new Mycobacterium leprae antigens to be used in methods and means for detection and \*\*\*diagnostics\*\*\* of M. leprae infections in subjects, in particular in the early stages of infection and in paucibacillary infections, which remain undetected using conventional \*\*\*diagnostic\*\*\* methods. The antigens disclosed in the invention are specific for M. leprae and the \*\*\*diagnostic\*\*\* method does not yield 'false pos.' results in individuals having an immune response against other Mycobacterial species, such as M. tuberculosis, M. bovis, M. \*\*\*paratuberculosis\*\*\*, M. avium, M. smegmatis,, M. ulcerans, M. microti, and M. marinum, or BCG vaccinated individuals.

Thus, using bioinformatic anal. the antigen genes ML0573, ML0574, ML0575, ML0576, ML1602, ML1603, ML1604, ML1788, ML1989, ML1990, ML2283 and ML2567 were found to be unique to *M. leprae*. It was demonstrated, that all of above genes were expressed at the mRNA level in human leprosy tissue. Paucibacillary and reactional leprosy patients and healthy household contacts of leprosy patients produced significant levels of \*\*\*interferon\*\*\* (IFN)-.gamma. in response to the five unique *M. leprae* antigens encoded by ML0576, ML1989, ML1990, ML2283 and ML2567. Provided are gene and protein sequences, as well as sequences for epitope peptides for *M. leprae*-specific antigens ML0576, ML1989, ML1990, ML2283 and ML2567. A method for identifying Mycobacterium leprae antigens is also provided.

TI Sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* *M. leprae*, particularly in the early stages and paucibacillary infections

AB The current invention discloses new Mycobacterium leprae antigens to be used in methods and means for detection and \*\*\*diagnostics\*\*\* of *M. leprae* infections in subjects, in particular in the early stages of infection and in paucibacillary infections, which remain undetected using conventional \*\*\*diagnostic\*\*\* methods. The antigens disclosed in the invention are specific for *M. leprae* and the \*\*\*diagnostic\*\*\* method does not yield 'false pos.' results in individuals having an immune response against other Mycobacterial species, such as *M. tuberculosis*, *M. bovis*, *M. paratuberculosis*, *M. avium*, *M. smegmatis*, *M. ulcerans*, *M. microti*, and *M. marinum*, or BCG vaccinated individuals. Thus, using bioinformatic anal. the . . . in human leprosy tissue. Paucibacillary and reactional leprosy patients and healthy household contacts of leprosy patients produced significant levels of \*\*\*interferon\*\*\* (IFN)-.gamma. in response to the five unique *M. leprae* antigens encoded by ML0576, ML1989, ML1990, ML2283 and ML2567. Provided are. . .

ST sequence Mycobacterium leprae antigen epitope \*\*\*diagnoses\*\*\* infection; leprosy immunodiagnosis Mycobacterium leprae antigen epitope; vaccine Mycobacterium leprae antigen epitope

IT Receptors  
 RL: BSU (Biological study, unclassified); BIOL (Biological study) (4-1BB, anti-4-1BB agonistic antibody as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* *M. leprae*, particularly in early stages and paucibacillary infections)

IT Human groups  
 (Brazilian patients; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* *M. leprae*, particularly in early stages and paucibacillary infections)

IT Genetic element  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (CpG island, CpG, as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* *M. leprae*, particularly in early stages and paucibacillary infections)

IT Histocompatibility antigens  
 RL: BSU (Biological study, unclassified); BIOL (Biological study) (HLA, class I, identifying T-cell epitopes for, using computer algorithms; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* *M. leprae*, particularly in early stages and paucibacillary infections)

IT Histocompatibility antigens  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)

(HLA, class II, identifying T-cell epitopes for, using computer algorithms; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Proteins  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (LAG3 (lymphocyte activation gene-3), sol., as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML0573, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML0574, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML0575, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML0576, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Antigens  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (ML0576; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1602, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1603, expressed in human leprosy tissue; sequences for Mycobacterium

leprae-specific antigens, and methods for treating and  
 \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and  
 paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study,  
 unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1604, expressed in human leprosy tissue; sequences for Mycobacterium  
 leprae-specific antigens, and methods for treating and  
 \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and  
 paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study,  
 unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1788, expressed in human leprosy tissue; sequences for Mycobacterium  
 leprae-specific antigens, and methods for treating and  
 \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and  
 paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study,  
 unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1989, expressed in human leprosy tissue; sequences for Mycobacterium  
 leprae-specific antigens, and methods for treating and  
 \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and  
 paucibacillary infections)

IT Antigens  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role,  
 unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use);  
 PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL  
 (Biological study); USES (Uses)  
 (ML1989; sequences for Mycobacterium leprae-specific antigens, and  
 methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly  
 in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study,  
 unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1990, expressed in human leprosy tissue; sequences for Mycobacterium  
 leprae-specific antigens, and methods for treating and  
 \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and  
 paucibacillary infections)

IT Antigens  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role,  
 unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use);  
 PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL  
 (Biological study); USES (Uses)  
 (ML1990; sequences for Mycobacterium leprae-specific antigens, and  
 methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly  
 in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study,  
 unclassified); PRP (Properties); BIOL (Biological study)  
 (ML2283, expressed in human leprosy tissue; sequences for Mycobacterium  
 leprae-specific antigens, and methods for treating and  
 \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and  
 paucibacillary infections)

IT Antigens  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role,  
 unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use);

PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (ML2283; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML2567, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Antigens  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (ML2567; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Lipopeptides  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (Pam3Cys, as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Immunostimulants  
 (adjuvants, DA/TDB; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Immunostimulants  
 (adjuvants, DDA/MPL; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Immunostimulants  
 (adjuvants; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Monocyte  
 (anal., in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Diagnostic\*\*\* agents  
 Vaccines  
 (antigens or epitopes as; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Lipid A  
 Lipopolysaccharides  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Mycobacterium  
 (as recombinant expression host; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and



paucibacillary infections)

IT Flagellins  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (bacterial, as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT CD40 (antigen)  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (binding CD40 ligand or antibody, as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Mammalia  
 ( \*\*\*diagnosis\*\*\* and therapy; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Mycobacterium avium  
 Mycobacterium bovis  
 Mycobacterium marinum  
 Mycobacterium microti  
 Mycobacterium smegmatis  
 Mycobacterium tuberculosis  
 Mycobacterium ulcerans  
 (differentiating from; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Leprosy  
 (early stages \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT T cell (lymphocyte)  
 (epitopes; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Epitopes  
 (from ML0576, ML1989, ML1990, ML2283 and ML2567 antigens; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT T cell (lymphocyte)  
 (helper cell, measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Algorithm  
 (identifying HLA class I and/or class II T-cell epitopes using; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Diagnosis\*\*\*  
 (immunodiagnosis, of ML0576, ML1989, ML1990, ML2283 and ML2567 antigens; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Blood analysis

(in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Interleukin\*\*\* 10  
     \*\*\*Interleukin\*\*\* 15  
     \*\*\*Interleukin\*\*\* 2  
     \*\*\*Interleukin\*\*\* 4  
     \*\*\*Interleukin\*\*\* 6

Macrophage inflammatory protein 1.beta.  
 Transforming growth factor .beta.  
 Tumor necrosis factors

RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
 (measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Antibodies and Immunoglobulins  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (monoclonal, anti-4-1BB, agonistic, as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Genome  
 (of M. leprae, identifying unique antigen gene candidates in; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Protein sequences  
 (of M. leprae-specific antigens ML0576, ML1989, ML1990, ML2283 and ML2567; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT DNA sequences  
 (of M. leprae-specific genes ML0576, ML1989, ML1990, ML2283 and ML2567; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Blood cell  
 (of infected subject, IFN-.gamma. response in; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Interleukin\*\*\* 12  
 RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
 (p70, measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Human  
 (patients; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Infection  
 (paucibacillary, \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and

\*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Bioinformatics  
(sequence annotation, M. leprae unique genes; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Molecular cloning  
Mycobacterium leprae  
Test kits  
(sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Skin  
(test, by applying antigen under top skin; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Mycobacterium BCG  
(vaccine, differentiating from; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Interferons\*\*\*  
RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
(.alpha., measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Interferons\*\*\*  
RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
(.beta., measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Interferons\*\*\*  
RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
(.gamma., measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 141256-04-4, QS21  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(MPL, as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 946442-88-2 946442-91-7  
RL: PRP (Properties)  
(Unclaimed; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in the early stages and paucibacillary infections)

IT 946400-78-8 946400-79-9 946400-80-2 946400-81-3 946400-82-4  
RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use);

PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (amino acid sequence, epitope; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 946442-52-0 946442-53-1 946442-54-2 946442-55-3 946442-56-4  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (amino acid sequence; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 24939-03-5, Poly(I:C) 87420-41-5, Pam3Cys 911642-39-2, IC 31  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 83869-56-1, GM-CSF  
 RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
 (measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 946442-57-5, DNA (Mycobacterium leprae gene ML0576) 946442-58-6, DNA (Mycobacterium leprae gene ML1989) 946442-59-7, DNA (Mycobacterium leprae gene ML1990) 946442-60-0, DNA (Mycobacterium leprae gene ML2283) 946442-61-1, DNA (Mycobacterium leprae gene ML2567)  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (nucleotide sequence; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 946442-98-4 946442-99-5 946443-00-1 946443-01-2 946443-02-3  
 946443-03-4 946443-04-5 946443-05-6 946443-06-7 946443-07-8  
 946443-08-9 946443-09-0  
 RL: PRP (Properties)  
 (unclaimed nucleotide sequence; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in the early stages and paucibacillary infections)

IT 946442-86-0 946442-87-1 946442-89-3 946442-90-6 946442-92-8  
 946442-93-9 946442-94-0 946442-95-1 946442-96-2 946442-97-3  
 RL: PRP (Properties)  
 (unclaimed protein sequence; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in the early stages and paucibacillary infections)

L10 ANSWER 2 OF 19 MEDLINE on STN  
 AN 2007416292 MEDLINE <<LOGINID::20080325>>  
 DN PubMed ID: 17502388  
 TI Influence of Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* on colitis development and specific immune responses during disease.  
 AU Singh Udai P; Singh Shailesh; Singh Rajesh; Karls Russell K; Quinn

Frederick D; Potter Morris E; Lillard James W Jr  
 CS Brown Cancer Center, Department of Microbiology and Immunology, University  
 of Louisville, 580 S. Preston Street, Baxter II/Room 304C, Louisville, KY  
 40202, USA.

NC AI 57808 (United States NIAID)  
 GM 08248 (United States NIGMS)  
 MD 000525 (United States NCMHD)  
 RR 03034 (United States NCRR)

SO Infection and immunity, (2007 Aug) Vol. 75, No. 8, pp. 3722-8. Electronic  
 Publication: 2007-05-14.  
 Journal code: 0246127. ISSN: 0019-9567.

CY United States

DT Journal; Article; (JOURNAL ARTICLE)  
 (RESEARCH SUPPORT, N.I.H., EXTRAMURAL)  
 (RESEARCH SUPPORT, NON-U.S. GOV'T)

LA English

FS Priority Journals

EM 200709

ED Entered STN: 20 Jul 2007  
 Last Updated on STN: 7 Sep 2007  
 Entered Medline: 6 Sep 2007

AB The granulomatous and intramural inflammation observed in cases of  
 inflammatory bowel diseases (IBD) and veterinary Johne's disease suggests  
 that Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* is a causative  
 agent. However, an incomplete understanding of the immunological steps  
 responsible for the pathologies of IBD makes this conclusion uncertain.  
 Sera from \*\*\*interleukin\*\*\* -10-deficient (IL-10(-/-)) mice with  
 spontaneous colitis displayed significantly higher M. avium subsp.  
 \*\*\*paratuberculosis\*\*\* -specific immunoglobulin G2a antibody responses  
 than did sera from similar mice without disease. Pathogen-free IL-10(-/-)  
 mice received control vehicle or the vehicle containing heat-killed or  
 live M. avium subsp. \*\*\*paratuberculosis\*\*\*. Mucosal CD4(+) T cells  
 from the mice that developed colitis proliferated and secreted higher  
 levels of gamma \*\*\*interferon\*\*\* and tumor necrosis factor alpha after  
 ex vivo stimulation with a Vbeta11(+) T-cell receptor-restricted peptide  
 from the MPT59 antigen (Ag85B) than those secreted from cells from mice  
 before the onset of colitis. The data from this study provide important  
 information regarding the mechanisms of colitis in IL-10(-/-) mice, which  
 are driven in part by Ag85B-specific T cells. The data suggest a  
 plausible mechanism of Ag-specific T-cell responses in colitis driven by  
 potent Ags conserved in Mycobacterium species.

TI Influence of Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* on  
 colitis development and specific immune responses during disease.

AB . . . and intramural inflammation observed in cases of inflammatory  
 bowel diseases (IBD) and veterinary Johne's disease suggests that  
 Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* is a causative  
 agent. However, an incomplete understanding of the immunological steps  
 responsible for the pathologies of IBD makes this conclusion uncertain.  
 Sera from \*\*\*interleukin\*\*\* -10-deficient (IL-10(-/-)) mice with  
 spontaneous colitis displayed significantly higher M. avium subsp.  
 \*\*\*paratuberculosis\*\*\* -specific immunoglobulin G2a antibody responses  
 than did sera from similar mice without disease. Pathogen-free IL-10(-/-)  
 mice received control vehicle or the vehicle containing heat-killed or  
 live M. avium subsp. \*\*\*paratuberculosis\*\*\*. Mucosal CD4(+) T cells  
 from the mice that developed colitis proliferated and secreted higher  
 levels of gamma \*\*\*interferon\*\*\* and tumor necrosis factor alpha after  
 ex vivo stimulation with a Vbeta11(+) T-cell receptor-restricted peptide

from the MPT59 antigen (Ag85B). . . .

CT . . . .

Antigens, Bacterial: IM, immunology

CD4-Positive T-Lymphocytes: IM, immunology

Colitis: IM, immunology

\*Colitis: MI, microbiology

\*Colitis: PA, pathology

Disease Models, Animal

\*\*\* Enzyme-Linked Immunosorbent Assay\*\*\*

Humans

Immunoglobulin G: BL, blood

\*\*\* Interferon Type II: BI, biosynthesis\*\*\*

\*\*\* Interleukin-10: DF, deficiency\*\*\*

Intestinal Mucosa: IM, immunology

Ligands

Mice

Mice, Knockout

\*\*\*\*Mycobacterium avium subsp. paratuberculosis: IM, immunology\*\*\*

\*\*\*\*Paratuberculosis: IM, immunology\*\*\*

\*\*\*\*Paratuberculosis: PA, pathology\*\*\*

Peptides: IM, immunology

Receptors, Antigen, T-Cell: IM, immunology

Receptors, CXCR3

Receptors, Chemokine: AG, agonists

Receptors, Chemokine: IM, immunology

. . . .

RN \*\*\*130068-27-8 (Interleukin-10)\*\*\* ; \*\*\*82115-62-6 (Interferon Type\*\*\*  
 \*\*\* II)\*\*\*

CN. . . 0 (Peptides); 0 (Receptors, Antigen, T-Cell); 0 (Receptors, CXCR3);  
 0 (Receptors, Chemokine); 0 (Tumor Necrosis Factor-alpha); 0 (antigen 85B,  
 Mycobacterium \*\*\*paratuberculosis\*\*\* )

L10 ANSWER 3 OF 19 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN

AN 2006:496945 BIOSIS <<LOGINID::20080325>>

DN PREV200600503265

TI Disturbed cytokine response to mycobacterium avium subspecies  
 \*\*\*paratuberculosis\*\*\* is dysregulated in patients with Crohn's  
 disease.

AU Sibartie, Shomik; Keohane, John; Scully, Paul; O'Neill, Shaun; O'Mahony,  
 Jim; O'Mahony, Liam; Shanahan, Fergus

SO Gastroenterology, (APR 2006) Vol. 130, No. 4, Suppl. 2, pp. A240.  
 Meeting Info.: Digestive Disease Week Meeting/107th Annual Meeting of the  
 American-Gastroenterological-Association. Los Angeles, CA, USA. May 19  
 -24, 2006. Amer Gastroenterol Assoc Inst.  
 CODEN: GASTAB. ISSN: 0016-5085.

DT Conference; (Meeting)

LA English

ED Entered STN: 4 Oct 2006

AB Background: Mycobacterium avium subspecies \*\*\*paratuberculosis\*\*\*  
 (MAP) has been a source of controversy since it was first suggested as a  
 possible cause for Crohn's disease. While a number of studies have  
 focused on identification of the organism in Crohn's disease tissues and  
 others have assessed the serologic response to MAP, few studies have  
 examined the cellular immune response to MAP. Aim: To compare the  
 cellular response to Mycobacterium avium subspecies

\*\*\*paratuberculosis\*\*\* between Crohn's disease patients and healthy volunteers. Methods: Peripheral blood mononuclear cells (PBMCs) were isolated from 24 Crohn's disease patients and 20 healthy volunteers. After in-vitro co-incubation for 72 hrs with MAP ATCC 43019 at several concentrations, supernatants were harvested and analysed for IL2, IL-4, IL-6, IL-8, IL-10, TNF-alpha and IFN-gamma using cytometric bead analysis. PBMCs stimulated with Salmonella typhimurium (ST) were used as positive controls. Results: Compared to healthy volunteers, PBMCs from Crohn's disease patients secreted higher levels ( $p < 0.05$ ) of IL-6 (4265 +/- 260 vs. 2865 +/- 386 pg/ml), TNF-alpha (2190 +/- 247 vs. 1200 +/- 228 pg/ml) and IL-10 (265 +/- 53 vs. 81 +/- 12 pg/ml) upon exposure to MAP (1 MAP bacterial cell:1 PBMC) but showed no difference when exposed to ST. There was a lower IFN-gamma response from Crohn's disease patients to MAP (322 +/- 125 vs. 1658 +/- 424 pg/ml) but this also occurred in response to ST. The ratio of IFN-gamma to IL-10 was significantly lower for Crohn's disease patients' PBMCs exposed to MAP compared to healthy volunteers ( $p < 0.05$ ) but not for ST. There were no differences in IL-8, IL-2 and IL-4 levels. Prior BCG vaccination and concurrent immunosuppressives had no impact on the levels of cytokines secreted. Conclusion: A diminished Th1 response to MAP in Crohn's disease patients may allow for prolonged intracellular survival of MAP in phagocytic cells. This might account for the increased frequency of MAP detection in patients with Crohn's disease but does not imply cause and effect.

TI Disturbed cytokine response to mycobacterium avium subspecies  
 \*\*\*paratuberculosis\*\*\* is dysregulated in patients with Crohn's disease.

AB Background: Mycobacterium avium subspecies \*\*\*paratuberculosis\*\*\* (MAP) has been a source of controversy since it was first suggested as a possible cause for Crohn's disease. While. . . few studies have examined the cellular immune response to MAP. Aim: To compare the cellular response to Mycobacterium avium subspecies  
 \*\*\*paratuberculosis\*\*\* between Crohn's disease patients and healthy volunteers. Methods: Peripheral blood mononuclear cells (PBMCs) were isolated from 24 Crohn's disease patients. . .

IT . . .  
 system, blood and lymphatics, PBMC; phagocytic cell: immune system

IT Diseases  
 Crohn's disease: digestive system disease, immune system disease, etiology, \*\*\*diagnosis\*\*\*

IT Chemicals & Biochemicals  
 cytokines; IFN-gamma [ \*\*\*interferon\*\*\* -gamma]; IL-10 [ \*\*\*interleukin\*\*\* -10]; TNF-alpha [tumor necrosis factor-alpha]; IL-6 [ \*\*\*interleukin\*\*\* -6]; IL-8 [ \*\*\*interleukin\*\*\* -8]; IL-2 [ \*\*\*interleukin\*\*\* -2]; IL-4 [ \*\*\*interleukin\*\*\* -4]

ORGN . . .  
 Primates, Vertebrates

ORGN Classifier  
 Mycobacteriaceae 08881  
 Super Taxa  
 Mycobacteria; Actinomycetes and Related Organisms; Eubacteria;  
 Bacteria; Microorganisms  
 Organism Name  
 Mycobacterium avium \*\*\*paratuberculosis\*\*\* (subspecies): pathogen  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms

AN 2006:423091 BIOSIS <<LOGINID::20080325>>  
DN PREV200600423340  
TI Differential expression of genes encoding CD30L and P-selectin in cattle with Johne's disease: Progress toward a \*\*\*diagnostic\*\*\* gene expression signature.  
AU Skovgaard, Kerstin [Reprint Author]; Grell, Susanne Nedergaard; Heegaard, Peter M. H.; Jungersen, Gregers; Pudrith, Chas B.; Coussens, Paul M.  
CS Danish Inst Food and Vet Res, Dept Vet Diagnost and Res, Bulowsvej 27, DK-1790 Copenhagen V, Denmark  
kis@dfvf.dk  
SO Veterinary Immunology and Immunopathology, (AUG 15 2006) Vol. 112, No. 3-4, pp. 210-224.  
CODEN: VIIMDS. ISSN: 0165-2427.  
DT Article  
LA English  
ED Entered STN: 23 Aug 2006  
Last Updated on STN: 23 Aug 2006  
AB Mycobacterium avium subspecies \*\*\*paratuberculosis\*\*\* (Mycobacterium \*\*\*paratuberculosis\*\*\*), the causative agent of \*\*\*paratuberculosis\*\*\* (paraTB) or Johne's disease in ruminants, is a health problem for the global cattle industry with significant economic losses related to decreased milk production and reduced fertility. Commonly paraTB in cattle is \*\*\*diagnosed\*\*\* by antibody detection by serum enzyme-linked immunosorbent \*\*\*assay\*\*\* (ELISA), by detection of the pathogen by cultivation of individual faecal samples, or by in vitro measurement of cell mediated immune responses using the IFN-gamma test. There is an ongoing need for developing new \*\*\*diagnostic\*\*\* approaches as all currently available \*\*\*diagnostic\*\*\* tests for paraTB may fail to detect sub-clinical infection. We used cDNA microarrays to simultaneously measure expression of over 1300 host genes to help identify a subset of gene expression changes that might provide a unique gene expression signature for paraTB infection. In the present study, non-stimulated leukocytes isolated from 10 sub-clinical paraTB infected cows were examined for genes being expressed at significantly different levels than in similar cells from control cows with the same herd background. We included cattle (Holstein) from two locations (Denmark and USA) for the microarray experiment. Our results indicate that expression profiles of at least 52 genes are different in leukocytes from M. \*\*\*paratuberculosis\*\*\* infected cattle compared to control cattle.

Gene expression differences were verified by quantitative real-time reverse transcriptase polymerase chain reactions (qRT-PCR) on the same group of cattle (Holstein) used for the microarray experiment. In order to assess the generality of the observed gene expression, a second and different group of cattle (Jersey) was also examined using qRT-PCR. Out of the seven genes selected for qRT-PCR, CD30 ligand (CD30L) and P-selectin were consistently differentially expressed in freshly isolated leukocytes from paraTB infected and control animals of both breeds of cattle. Although further work is clearly needed to develop a more complete gene expression signature specific for paraTB, our results demonstrate that a subset of genes in leukocytes are consistently expressed at different levels, depending upon M. \*\*\*paratuberculosis\*\*\* infection status. (c) 2006 Elsevier B.V. All rights reserved.

TI Differential expression of genes encoding CD30L and P-selectin in cattle with Johne's disease: Progress toward a \*\*\*diagnostic\*\*\* gene expression signature.



AB Mycobacterium avium subspecies \*\*\*paratuberculosis\*\*\* (Mycobacterium  
 \*\*\*paratuberculosis\*\*\* ), the causative agent of  
 \*\*\*paratuberculosis\*\*\*  
 (paraTB) or Johne's disease in ruminants, is a health problem for the  
 global cattle industry with significant economic losses related to  
 decreased milk production and reduced fertility. Commonly paraTB in  
 cattle is \*\*\*diagnosed\*\*\* by antibody detection by serum enzyme-linked  
 immunosorbent \*\*\*assay\*\*\* (ELISA), by detection of the pathogen by  
 cultivation of individual faecal samples, or by in vitro measurement of  
 cell mediated immune responses using the IFN-gamma test. There is an  
 ongoing need for developing new \*\*\*diagnostic\*\*\* approaches as all  
 currently available \*\*\*diagnostic\*\*\* tests for paraTB may fail to  
 detect sub-clinical infection. We used cDNA microarrays to simultaneously  
 measure expression of over 1300. . . the microarray experiment. Our  
 results indicate that expression profiles of at least 52 genes are  
 different in leukocytes from M. \*\*\*paratuberculosis\*\*\* infected cattle  
 compared to control cattle. Gene expression differences were verified by  
 quantitative real-time reverse transcriptase polymerase chain reactions  
 (qRT-PCR). . . paraTB, our results demonstrate that a subset of genes  
 in leukocytes are consistently expressed at different levels, depending  
 upon M. \*\*\*paratuberculosis\*\*\* infection status. (c) 2006 Elsevier  
 B.V. All rights reserved.

IT . . .  
 Organisms  
 feces: digestive system; leukocyte: immune system, blood and lymphatics

IT Diseases  
 Johne's disease: bacterial disease, infectious disease

IT Diseases  
 \*\*\*paratuberculosis\*\*\* : bacterial disease, infectious disease,  
 etiology  
 \*\*\*Paratuberculosis\*\*\* (MeSH)

IT Chemicals & Biochemicals  
 IFN-gamma [ \*\*\*interferon\*\*\* -gamma]; cDNA [complementary DNA]

GEN. . . leukemia inhibitory factor mRNA gene] (Bovidae); bovine  
 TNF-alpha-CE gene [bovine tumor necrosis factor-alpha-converting enzyme  
 gene] (Bovidae); bovine IL-1RA gene [bovine \*\*\*interleukin\*\*\* -1  
 receptor antagonist mRNA gene] (Bovidae); bovine P-selectin gene [bovine  
 P-selectin mRNA gene] (Bovidae); bovine Caspase-7 gene [bovine Mch-7  
 isoform alpha. . .

L10 ANSWER 5 OF 19 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
 DUPLICATE 1

AN 2005:498376 BIOSIS <<LOGINID::20080325>>

DN PREV200510279086

TI Bovine NK cells can produce gamma \*\*\*interferon\*\*\* in response to the  
 secreted mycobacterial proteins ESAT-6 and MPP14 but not in response to  
 MPB70.

AU Olsen, Ingrid [Reprint Author]; Boysen, Preben; Kulberg, Siri; Hope, Jayne  
 C.; Jungersen, Gregers; Storset, Anne K.

CS Natl Vet Inst, POB 8156 DEP, N-0033 Oslo, Norway  
 Ingrid.Olsen@vetinst.no

SO Infection and Immunity, (SEP 2005) Vol. 73, No. 9, pp. 5628-5635.  
 CODEN: INFIBR. ISSN: 0019-9567.

DT Article

LA English

ED Entered STN: 16 Nov 2005  
 Last Updated on STN: 16 Nov 2005

AB Bovine NK cells have recently been characterized and the present study describes the interaction between NK cells, antigen-presenting cells, and secreted mycobacterial proteins. Gamma \*\*\*interferon\*\*\* (IFN-gamma) production by NK cells was seen in approximately 30% of noninfected calves in response to the Mycobacterium tuberculosis complex-specific protein ESAT-6, MPP14 from Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\*, and purified protein derivative (PPD) from M. tuberculosis. In contrast, no response was induced by MPB70, which is another M. tuberculosis complex-specific secreted antigen. The production of IFN-gamma by NK cells in whole blood in response to ESAT-6 and MPP14 was demonstrated using intracellular staining together with surface labeling for the NK cell-specific receptor, NKp46, or CD3. Furthermore, the depletion of NK cells from peripheral blood mononuclear cells completely abolished the IFN-gamma production. The response was mediated through stimulation of adherent cells and was largely independent of contact between adherent cells and the NK cells. Neutralization of \*\*\*interleukin\*\*\* -12 only partly inhibited IFN-gamma production, showing that other cytokines were also involved. The demonstration of NK cell-mediated IFN-gamma production in young cattle provides an explanation for the nonspecific IFN-gamma response frequently encountered in young cattle when using the IFN-gamma test in \*\*\*diagnosis\*\*\* of mycobacterial infections.

TI Bovine NK cells can produce gamma \*\*\*interferon\*\*\* in response to the secreted mycobacterial proteins ESAT-6 and MPP14 but not in response to MPB70.

AB. . . recently been characterized and the present study describes the interaction between NK cells, antigen-presenting cells, and secreted mycobacterial proteins. Gamma \*\*\*interferon\*\*\* (IFN-gamma) production by NK cells was seen in approximately 30% of noninfected calves in response to the Mycobacterium tuberculosis complex-specific protein ESAT-6, MPP14 from Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\*, and purified protein derivative (PPD) from M. tuberculosis. In contrast, no response was induced by MPB70, which is another M. . . . through stimulation of adherent cells and was largely independent of contact between adherent cells and the NK cells. Neutralization of \*\*\*interleukin\*\*\* -12 only partly inhibited IFN-gamma production, showing that other cytokines were also involved. The demonstration of NK cell-mediated IFN-gamma production in. . . cattle provides an explanation for the nonspecific IFN-gamma response frequently encountered in young cattle when using the IFN-gamma test in \*\*\*diagnosis\*\*\* of mycobacterial infections.

IT . . . cell; peripheral blood mononuclear cell: immune system, blood and lymphatics; antigen-presenting cell: immune system

IT Diseases mycobacterial infection: bacterial disease, \*\*\*diagnosis\*\*\* Mycobacterium Infections (MeSH)

IT Chemicals & Biochemicals CD3; \*\*\*interferon\*\*\* -gamma [IFN-gamma]; \*\*\*interleukin\*\*\* -12; ESAT-6; NKp46; purified protein derivative; MPB70; mycobacterial proteins; MPP14

ORGN . . .

Classifier Mycobacteriaceae 08881 Super Taxa Mycobacteria; Actinomycetes and Related Organisms; Eubacteria; Bacteria; Microorganisms

Organism Name

Mycobacterium tuberculosis (species): pathogen

Mycobacterium avium \*\*\*paratuberculosis\*\*\* (subspecies): pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

L10 ANSWER 6 OF 19 MEDLINE on STN  
AN 2005487701 MEDLINE <<LOGINID::20080325>>  
DN PubMed ID: 15992970  
TI Vaccination of sheep against M. \*\*\*paratuberculosis\*\*\* : immune parameters and protective efficacy.  
AU Begg D J; Griffin J F T  
CS Disease Research Laboratory, Department of Microbiology and Immunology, University of Otago, P.O. Box 56, Dunedin, New Zealand.  
SO Vaccine, (2005 Oct 10) Vol. 23, No. 42, pp. 4999-5008.  
Journal code: 8406899. ISSN: 0264-410X.  
CY Netherlands  
DT (CLINICAL TRIAL)  
Journal; Article; (JOURNAL ARTICLE)  
(RESEARCH SUPPORT, NON-U.S. GOV'T)  
LA English  
FS Priority Journals  
EM 200511  
ED Entered STN: 14 Sep 2005  
Last Updated on STN: 11 Nov 2005  
Entered Medline: 10 Nov 2005  
AB Johne's disease in ruminants is caused by the pathogenic bacterium Mycobacterium avium subspecies \*\*\*paratuberculosis\*\*\* (Map). Currently available Map commercial vaccines protect against clinical disease but not infection. In this study, the proprietary Johne's vaccine Neoparasec and an aqueous formulation of Map 316F (AquaVax) were tested in sheep. Detailed immunological examination of blood and gut-associated lymphoid tissues was carried out on animals after vaccination and challenge with virulent Map to identify markers of protective immunity. Neoparasec vaccination provided significant protection against disease while AquaVax did not. Immune animals had stronger cell-mediated responses and altered proportions of CD4+, CD8+, CD25+ and B cells in blood, spleen and the gut lymphatics, than diseased animals.  
TI Vaccination of sheep against M. \*\*\*paratuberculosis\*\*\* : immune parameters and protective efficacy.  
AB Johne's disease in ruminants is caused by the pathogenic bacterium Mycobacterium avium subspecies \*\*\*paratuberculosis\*\*\* (Map). Currently available Map commercial vaccines protect against clinical disease but not infection. In this study, the proprietary Johne's vaccine. . .  
CT . . .  
Vaccines: AD, administration & dosage  
\*Bacterial Vaccines: IM, immunology  
CD4-Positive T-Lymphocytes: IM, immunology  
CD8-Positive T-Lymphocytes: IM, immunology  
Disease Models, Animal  
\*\*\* Enzyme-Linked Immunosorbent Assay\*\*\*  
\*\*\* Interferon Type II: AN, analysis\*\*\*  
Intestinal Mucosa: IM, immunology  
Lymphocyte Activation  
Lymphocyte Subsets: IM, immunology  
\*\*\*\*Mycobacterium avium subsp. paratuberculosis: IM, immunology\*\*\*

\*\*\*\*Paratuberculosis: PC, prevention & control\*\*\*  
 \*\*\* Receptors, Interleukin-2: AN, analysis\*\*\*  
 Sheep  
 \*Sheep Diseases: PC, prevention & control  
 Spleen: IM, immunology  
 RN \*\*\*\*82115-62-6 (Interferon Type II)\*\*\*  
 CN 0 (Antibodies, Bacterial); 0 (Bacterial Vaccines); 0 (Receptors,  
 \*\*\*\*Interleukin\*\*\* -2)

L10 ANSWER 7 OF 19 CABA COPYRIGHT 2008 CABI on STN DUPLICATE 2  
 AN 2006:49179 CABA <<LOGINID::20080325>>  
 DN 20063031176  
 TI Inflammatory cytokine gene expression in different types of granulomatous  
 lesions during asymptomatic stages of bovine \*\*\*\*paratuberculosis\*\*\*  
 AU Tanaka, S.; Sato, M.; Onitsuka, T.; Kamata, H.; Yokomizo, Y.  
 CS Comparative Pathology Section, Kyushu Research Station, National Institute  
 of Animal Health, Chuzan-cho 2702, Kagoshima 891-0105, Japan.  
 tanakas@affrc.go.jp  
 SO Veterinary Pathology, (2005) Vol. 42, No. 5, pp. 579-588. 41 ref.  
 Publisher: American College of Veterinary Pathologists Inc. Lawrence  
 ISSN: 0300-9858  
 DOI: 10.1354/vp.42-5-579  
 CY United States  
 DT Journal  
 LA English  
 ED Entered STN: 2 Mar 2006  
 Last Updated on STN: 2 Mar 2006  
 AB The granulomatous lesions in bovine \*\*\*\*paratuberculosis\*\*\* have been  
 classified into two types, i.e., the lepromatous type and the tuberculoid  
 type. To clarify the immunopathologic mechanisms at the site of infection,  
 we compared inflammatory cytokine gene expression between the two types of  
 lesions. Samples were obtained from noninfected control cows (n=5) and  
 naturally infected cows (n=7) that were \*\*\*\*diagnosed\*\*\* by  
 enzyme-linked immunosorbent \*\*\*\*assay\*\*\* (ELISA) and faecal culture  
 test. Although none of the infected cows showed clinical signs,  
 tuberculoid lesions were observed in five cows (tuberculoid group) and  
 lepromatous lesions in two cows (lepromatous group). Among the cytokines  
 examined by reverse transcription-polymerase chain reaction (RT-PCR),  
 Th2-type cytokines \*\*\*\*interleukin\*\*\* -4 (IL-4) and IL-10, and Th1-type  
 cytokine IL-2 were expressed more significantly in the lepromatous group  
 than in the tuberculoid (P<0.01) and noninfected groups (P<0.05). No  
 statistical differences were observed in the expression of  
 \*\*\*\*interferon\*\*\* -gamma, IL-1 beta, TNF-alpha, and GM-CSF among  
 lepromatous, tuberculoid, and noninfected groups. Expression of  
 proinflammatory cytokine IL-12 mRNA, however, did not differ among the  
 three groups; IL-18 was expressed at lower levels in the lepromatous group  
 than in the tuberculoid group and the noninfected group (P<0.0001).  
 Moreover, the number of cells in which IL-18 mRNAs were detected by in  
 situ hybridization was markedly decreased in the lepromatous group. These  
 results indicate that the formation of lepromatous-type lesions or  
 tuberculoid-type lesions may be influenced by alterations in Th1/Th2-type  
 cytokine production and that IL-18 may play an important role in a  
 Th1-to-Th2 switch in \*\*\*\*paratuberculosis\*\*\* .  
 TI Inflammatory cytokine gene expression in different types of granulomatous  
 lesions during asymptomatic stages of bovine \*\*\*\*paratuberculosis\*\*\* .  
 AB The granulomatous lesions in bovine \*\*\*\*paratuberculosis\*\*\* have been  
 classified into two types, i.e., the lepromatous type and the tuberculoid

type. To clarify the immunopathologic mechanisms at. . . the two types of lesions. Samples were obtained from noninfected control cows (n=5) and naturally infected cows (n=7) that were \*\*\*diagnosed\*\*\* by enzyme-linked immunosorbent \*\*\*assay\*\*\* (ELISA) and faecal culture test. Although none of the infected cows showed clinical signs, tuberculoid lesions were observed in five. . . and lepromatous lesions in two cows (lepromatous group). Among the cytokines examined by reverse transcription-polymerase chain reaction (RT-PCR), Th2-type cytokines \*\*\*interleukin\*\*\* -4 (IL-4) and IL-10, and Th1-type cytokine IL-2 were expressed more significantly in the lepromatous group than in the tuberculoid (P<0.01) and noninfected groups (P<0.05). No statistical differences were observed in the expression of \*\*\*interferon\*\*\* -gamma, IL-1 beta, TNF-alpha, and GM-CSF among lepromatous, tuberculoid, and noninfected groups. Expression of proinflammatory cytokine IL-12 mRNA, however, did not. . . influenced by alterations in Th1/Th2-type cytokine production and that IL-18 may play an important role in a Th1-to-Th2 switch in \*\*\*paratuberculosis\*\*\* .

CT cows; cytokines; disease course; gene expression; genes; granuloma; histopathology; immunopathology; \*\*\*interferon\*\*\* ; \*\*\*interleukin\*\*\* 1; \*\*\*interleukin\*\*\* 10; \*\*\*interleukin\*\*\* 2; \*\*\*interleukin\*\*\* 4; \*\*\*interleukins\*\*\* ; messenger RNA; \*\*\*paratuberculosis\*\*\* ; tumour necrosis factor

ST \*\*\*interleukin\*\*\* 18

ORGN cattle; Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\*

L10 ANSWER 8 OF 19 CAPLUS COPYRIGHT 2008 ACS on STN DUPLICATE 3

AN 2004:1056118 CAPLUS <<LOGINID::20080325>>

DN 142:73022

TI Analysis of the immune response to Mycobacterium avium subsp. \*\*\*Paratuberculosis\*\*\* in experimentally infected calves

AU Koo, Hye Cheong; Park, Yong Ho; Hamilton, Mary Jo; Barrington, George M.; Davies, Christopher J.; Kim, Jong Bae; Dahl, John L.; Waters, W. Ray; Davis, William C.

CS Department of Veterinary Microbiology, College of Veterinary Medicine and School of Agricultural Biotechnology, Seoul National University, Seoul, S. Korea

SO Infection and Immunity (2004), 72(12), 6870-6883  
CODEN: INFIBR; ISSN: 0019-9567

PB American Society for Microbiology

DT Journal

LA English

AB Johne's disease of cattle is widespread and causes significant economic loss to producers. Control has been hindered by limited understanding of the immune response to the causative agent, Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* , and lack of an effective vaccine and sensitive specific \*\*\*diagnostic\*\*\* \*\*\*assays\*\*\* . The present study was conducted to gain insight into factors affecting the immune response to M. avium subsp. \*\*\*paratuberculosis\*\*\* . A persistent proliferative response to M. avium subsp. \*\*\*paratuberculosis\*\*\* purified protein deriv. and sol. M. avium subsp. \*\*\*paratuberculosis\*\*\* antigens was detected in orally infected neonatal calves 6 mo postinfection (p.i.) by flow cytometry (FC). CD4+ T cells with a memory phenotype (CD45R0+) expressing CD25 and CD26 were the predominant cell type responding to antigens. Few CD8+ T cells proliferated in response to antigens until 18 mo p.i. .gamma..delta. T cells did not appear to respond to antigen until 18 mo p.i. The majority of WC1- CD2- and a few WC1- CD2+ .gamma..delta. T cells expressed CD25 at time zero. By 18 mo, however, subsets of

.gamma..delta. T cells from both control and infected animals showed an increase in expression of CD25, ACT2, and CD26 in the presence of the antigens. Two populations of CD3-. non-T non-B null cells, CD2+ and CD2-, proliferated in cell cultures from some control and infected animals during the study, with and without antigen. The studies clearly show multicolor FC offers a consistent reliable way to monitor the evolution and changes in the immune response to M. avium subsp.

\*\*\*paratuberculosis\*\*\* that occur during disease progression.

RE.CNT 65 THERE ARE 65 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

TI Analysis of the immune response to Mycobacterium avium subsp.  
\*\*\*Paratuberculosis\*\*\* in experimentally infected calves

AB . . . to producers. Control has been hindered by limited understanding of the immune response to the causative agent, Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\*, and lack of an effective vaccine and sensitive specific \*\*\*diagnostic\*\*\* \*\*\*assays\*\*\*. The present study was conducted to gain insight into factors affecting the immune response to M. avium subsp. \*\*\*paratuberculosis\*\*\*. A persistent proliferative response to M. avium subsp. \*\*\*paratuberculosis\*\*\* purified protein deriv. and sol. M. avium subsp. \*\*\*paratuberculosis\*\*\* antigens was detected in orally infected neonatal calves 6 mo postinfection (p.i.) by flow cytometry (FC). CD4+ T cells with. . . FC offers a consistent reliable way to monitor the evolution and changes in the immune response to M. avium subsp. \*\*\*paratuberculosis\*\*\* that occur during disease progression.

ST Mycobacterium infection cattle \*\*\*interleukin\*\*\* \*\*\*interferon\*\*\*  
TCR flow cytometry

IT \*\*\*Interleukin\*\*\* 2 receptors  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(.alpha. chain; flow cytometric anal. of immune response to Mycobacterium avium infection in calves)

IT \*\*\*Interferons\*\*\*  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(.gamma.; flow cytometric anal. of immune response to Mycobacterium avium infection in calves)

L10 ANSWER 9 OF 19 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
DUPLICATE 4

AN 2004:178760 BIOSIS <<LOGINID::20080325>>

DN PREV200400179647

TI Cytokine gene expression in peripheral blood mononuclear cells and tissues of cattle infected with Mycobacterium avium subsp.  
\*\*\*paratuberculosis\*\*\* : Evidence for an inherent proinflammatory gene expression pattern.

AU Coussens, Paul M. [Reprint Author]; Verman, Nitin; Coussens, Marc A.; Elftman, Michael D.; McNulty, Amanda M.

CS Department of Animal Science, Michigan State University, 1205H Anthony Hall, East Lansing, MI, 48824, USA  
coussens@msu.edu

SO Infection and Immunity, (March 2004) Vol. 72, No. 3, pp. 1409-1422. print. ISSN: 0019-9567 (ISSN print).

DT Article

LA English

ED Entered STN: 31 Mar 2004  
Last Updated on STN: 31 Mar 2004

AB In cattle and other ruminants, infection with the intracellular pathogen Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* results in a

granulomatous enteritis (Johne's disease) that is often fatal. The key features of host immunity to *M. avium* subsp. **\*\*\*paratuberculosis\*\*\*** infection include an appropriate early proinflammatory and cytotoxic response (Th1-like) that eventually gives way to a predominant antibody-based response (Th2-like). Clinical disease symptoms often appear subsequent to waning of the Th1-like immune response. Understanding why this shift in the immune response occurs and the underlying molecular mechanisms involved is critical to future control measures and **\*\*\*diagnosis\*\*\***. Previous studies have suggested that *M. avium* subsp. **\*\*\*paratuberculosis\*\*\*** may suppress gene expression in peripheral blood mononuclear cells (PBMCs) from infected cows, despite a continued inflammatory reaction at sites of infection. In the present study, we tested the hypothesis that exposure to *M. avium* subsp.

**\*\*\*paratuberculosis\*\*\*** suppresses a proinflammatory gene expression pattern in PBMCs from infected cows. To do this, we examined expression of genes encoding **\*\*\*interleukin\*\*\*** -1alpha (IL-1alpha), IL-2, IL-4, IL-5, IL-6, IL-8, IL-10, IL-12p35, IL-16, and IL-18, as well as genes encoding gamma **\*\*\*interferon\*\*\*** (IFN-gamma), transforming growth factor beta (TGF-beta), and tumor necrosis factor alpha (TNF-alpha), in PBMCs, intestinal lesions, and mesenteric lymph nodes of cattle naturally infected with *M. avium* subsp. **\*\*\*paratuberculosis\*\*\***. Cytokine gene expression in these cells and tissues was compared to expression in similar cells and tissues from control uninfected cattle. Our comprehensive results demonstrate that for most cytokine genes, including the genes encoding IFN-gamma, TGF-beta, TNF-alpha, IL-1alpha, IL-4, IL-6, IL-8, and IL-12p35, differential expression in PBMCs from infected and control cattle did not require stimulation with *M. avium* subsp.

**\*\*\*paratuberculosis\*\*\***. In fact, stimulation with *M. avium* subsp.

**\*\*\*paratuberculosis\*\*\*** tended to reduce the differential expression observed in infected and uninfected cows for genes encoding IFN-gamma, IL-1alpha, and IL-6. Only IL-10 gene expression was consistently enhanced by *M. avium* subsp. **\*\*\*paratuberculosis\*\*\*** stimulation of PBMCs from subclinically infected cattle. In ileal tissues from *M. avium* subsp.

**\*\*\*paratuberculosis\*\*\*** -infected cattle, expression of the genes encoding IFN-gamma, TGF-beta, IL-5, and IL-8 was greater than the expression in comparable tissues from control uninfected cattle, while expression of the gene encoding IL-16 was lower in tissues from infected cattle than in control tissues. Mesenteric lymph nodes draining sites of *M. avium* subsp.

**\*\*\*paratuberculosis\*\*\*** infection expressed higher levels of IL-1alpha, IL-8, IL-2, and IL-10 mRNA than similar tissues from control uninfected cattle expressed. In contrast, the genes encoding TGF-beta and IL-16 were expressed at lower levels in lymph nodes from infected cattle than in tissues from uninfected cattle. Taken together, our results suggest that cells or other mechanisms capable of limiting proinflammatory responses to *M. avium* subsp. **\*\*\*paratuberculosis\*\*\*** develop in infected cattle and that a likely place for development and expansion of these cell populations is the mesenteric lymph nodes draining sites of infection.

TI Cytokine gene expression in peripheral blood mononuclear cells and tissues of cattle infected with *Mycobacterium avium* subsp.

**\*\*\*paratuberculosis\*\*\*** : Evidence for an inherent proinflammatory gene expression pattern.

AB In cattle and other ruminants, infection with the intracellular pathogen *Mycobacterium avium* subsp. **\*\*\*paratuberculosis\*\*\*** results in a granulomatous enteritis (Johne's disease) that is often fatal. The key features of host immunity to *M. avium* subsp. **\*\*\*paratuberculosis\*\*\*** infection include an appropriate early proinflammatory and cytotoxic

response (Th1-like) that eventually gives way to a predominant antibody-based response (Th2-like).. . . this shift in the immune response occurs and the underlying molecular mechanisms involved is critical to future control measures and \*\*\*diagnosis\*\*\* . Previous studies have suggested that *M. avium* subsp. \*\*\*paratuberculosis\*\*\* may suppress gene expression in peripheral blood mononuclear cells (PBMCs) from infected cows, despite a continued inflammatory reaction at sites of infection. In the present study, we tested the hypothesis that exposure to *M. avium* subsp. \*\*\*paratuberculosis\*\*\* suppresses a proinflammatory gene expression pattern in PBMCs from infected cows. To do this, we examined expression of genes encoding \*\*\*interleukin\*\*\* -1alpha (IL-1alpha), IL-2, IL-4, IL-5, IL-6, IL-8, IL-10, IL-12p35, IL-16, and IL-18, as well as genes encoding gamma \*\*\*interferon\*\*\* (IFN-gamma), transforming growth factor beta (TGF-beta), and tumor necrosis factor alpha (TNF-alpha), in PBMCs, intestinal lesions, and mesenteric lymph nodes of cattle naturally infected with *M. avium* subsp.

\*\*\*paratuberculosis\*\*\* . Cytokine gene expression in these cells and tissues was compared to expression in similar cells and tissues from control uninfected. . . IL-8, and IL-12p35, differential expression in PBMCs from infected and control cattle did not require stimulation with *M. avium* subsp. \*\*\*paratuberculosis\*\*\* . In fact, stimulation with *M. avium* subsp. \*\*\*paratuberculosis\*\*\* tended to reduce the differential expression observed in infected and uninfected cows for genes encoding IFN-gamma, IL-1alpha, and IL-6. Only IL-10 gene expression was consistently enhanced by *M. avium* subsp. \*\*\*paratuberculosis\*\*\* stimulation of PBMCs from subclinically infected cattle. In ileal tissues from *M. avium* subsp. \*\*\*paratuberculosis\*\*\* -infected cattle, expression of the genes encoding IFN-gamma, TGF-beta, IL-5, and IL-8 was greater than the expression in comparable tissues from. . . was lower in tissues from infected cattle than in control tissues. Mesenteric lymph nodes draining sites of *M. avium* subsp. \*\*\*paratuberculosis\*\*\* infection expressed higher levels of IL-1alpha, IL-8, IL-2, and IL-10 mRNA than similar tissues from control uninfected cattle expressed. In. . . cattle. Taken together, our results suggest that cells or other mechanisms capable of limiting proinflammatory responses to *M. avium* subsp. \*\*\*paratuberculosis\*\*\* develop in infected cattle and that a likely place for development and expansion of these cell populations is the mesenteric. . .

```
IT . . .
    lymph node: blood and lymphatics, digestive system, immune system;
    peripheral blood mononuclear cell: blood and lymphatics, immune system
IT Diseases
    ***paratuberculosis*** : bacterial disease, infectious disease,
    genetics, immunology, Johne's disease
    ***Paratuberculosis*** (MeSH)
IT Chemicals & Biochemicals
    proinflammatory genes: expression pattern
ORGN . . .
    Vertebrates
ORGN Classifier
    Mycobacteriaceae 08881
    Super Taxa
    Mycobacteria; Actinomycetes and Related Organisms; Eubacteria;
    Bacteria; Microorganisms
Organism Name
    Mycobacterium avium ssp. ***paratuberculosis*** (subspecies):
    pathogen
```



# Taxa Notes

Bacteria, Eubacteria, Microorganisms

GEN cattle IFN-gamma gene [cattle \*\*\*interferon\*\*\* -gamma gene] (Bovidae);  
cattle IL-1-alpha gene [cattle \*\*\*interleukin\*\*\* -1-alpha gene]  
(Bovidae); cattle IL-10 gene [cattle \*\*\*interleukin\*\*\* -10 gene]  
(Bovidae); cattle IL-12p35 gene [cattle \*\*\*interleukin\*\*\* -12p35 gene]  
(Bovidae); cattle IL-16 gene [cattle \*\*\*interleukin\*\*\* -16 gene]  
(Bovidae); cattle IL-18 gene [cattle \*\*\*interleukin\*\*\* -18 gene]  
(Bovidae); cattle IL-2 gene [cattle \*\*\*interleukin\*\*\* -2 gene]  
(Bovidae); cattle IL-4 gene [cattle \*\*\*interleukin\*\*\* -4 gene]  
(Bovidae); cattle IL-5 gene [cattle \*\*\*interleukin\*\*\* -5 gene]  
(Bovidae); cattle IL-6 gene [cattle \*\*\*interleukin\*\*\* -6 gene]  
(Bovidae); cattle IL-8 gene [cattle \*\*\*interleukin\*\*\* -8 gene]  
(Bovidae); cattle TGF-beta gene [cattle transforming growth factor-beta  
gene] (Bovidae); cattle TNF-alpha gene [cattle tumor necrosis factor-alpha  
gene] (Bovidae)

L10 ANSWER 10 OF 19 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2004:885718 CAPLUS <<LOGINID::20080325>>

DN 141:363746

TI Development of early-stage \*\*\*diagnostic\*\*\* method for Johne disease  
by using anti-IL-10 antibody

AU Momotani, Eiichi; Mori, Yasuyuki

CS Natl. Agric. Bio-oriented Res. Org., Natl. Inst. Animal Health, Tsukuba,  
305-0856, Japan

SO BRAIN Techno News (2004), 105, 18-24

CODEN: BTEEEC; ISSN: 1345-5958

PB Nogyo, Seibutsukei Tokutei Sangyo Gijutsu Kenkyu Kiko, Seibutsukei Tokutei  
Sangyo Gijutsu Kenkyu Shien Senta

DT Journal; General Review

LA Japanese

AB A review on early-stage \*\*\*diagnosis\*\*\* of Johne's disease (  
\*\*\*paratuberculosis\*\*\* ) in cattle by modified \*\*\*interferon\*\*\*  
.gamma. ELISA \*\*\*assay\*\*\* using IL-10 neutralizing antibody, and its  
effectiveness.

TI Development of early-stage \*\*\*diagnostic\*\*\* method for Johne disease  
by using anti-IL-10 antibody

AB A review on early-stage \*\*\*diagnosis\*\*\* of Johne's disease (  
\*\*\*paratuberculosis\*\*\* ) in cattle by modified \*\*\*interferon\*\*\*  
.gamma. ELISA \*\*\*assay\*\*\* using IL-10 neutralizing antibody, and its  
effectiveness.

ST review cattle Johne disease \*\*\*diagnosis\*\*\* ELISA \*\*\*interleukin\*\*\*  
10 antibody; \*\*\*paratuberculosis\*\*\* cattle \*\*\*diagnosis\*\*\*  
\*\*\*interferon\*\*\* gamma ELISA review

IT Bos taurus

Mycobacterium avium \*\*\*paratuberculosis\*\*\*

(early-stage \*\*\*diagnosis\*\*\* method for Johne's disease using  
anti-IL-10 antibody)

IT \*\*\*Interleukin\*\*\* 10

RL: BSU (Biological study, unclassified); BIOL (Biological study)

(early-stage \*\*\*diagnosis\*\*\* method for Johne's disease using  
anti-IL-10 antibody)

IT Immunoassay

(enzyme-liked immunosorbent \*\*\*assay\*\*\* ; early-stage

\*\*\*diagnosis\*\*\* method for Johne's disease using anti-IL-10

antibody)

IT \*\*\*Diagnosis\*\*\*

(immunodiagnosis; early-stage \*\*\*diagnosis\*\*\* method for John's disease using anti-IL-10 antibody)

IT Infection  
 ( \*\*\*paratuberculosis\*\*\* , John's disease; early-stage \*\*\*diagnosis\*\*\* method for John's disease using anti-IL-10 antibody)

IT Antibodies and Immunoglobulins  
 RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (to IL-10; early-stage \*\*\*diagnosis\*\*\* method for John's disease using anti-IL-10 antibody)

IT \*\*\*Interferons\*\*\*  
 RL: ANT (Analyte); BSU (Biological study, unclassified); DGN (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (.gamma.; early-stage \*\*\*diagnosis\*\*\* method for John's disease using anti-IL-10 antibody)

L10 ANSWER 11 OF 19 CAPLUS COPYRIGHT 2008 ACS on STN  
 AN 2003:472526 CAPLUS <<LOGINID::20080325>>  
 DN 139:30816  
 TI Peptide T and analogs thereof for the stimulation of cytotoxic T lymphocyte (CTL) responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use  
 IN Ruscetti, Francis W.; Ruff, Michael R.  
 PA The Government of the United States of America, as Represented by the Secretary Department of Health and Human Services National Institutes of Health, USA  
 SO PCT Int. Appl., 43 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2003050136	A2	20030619	WO 2002-US39109	20021206
	WO 2003050136	A3	20031204		
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
	AU 2002357093	A1	20030623	AU 2002-357093	20021206
PRAI	US 2001-338971P	P	20011207		
	WO 2002-US39109	W	20021206		
OS	MARPAT 139:30816				
AB	The invention provides a method of increasing cytotoxic T lymphocyte (CTL) activity in a subject, comprising administering a CTL activity-stimulating amt. of peptide T or an analog thereof. A method of increasing .gamma.-***interferon*** (IFN-.gamma.) secretion in a subject comprises administering an IFN-.gamma. secretion-increasing amt. of peptide T or an analog thereof. A method of increasing ***interleukin*** 2 (IL-2)				

secretion in a subject comprises administering an IL-2 secretion-increasing amt. of peptide T or an analog thereof. A method of treating a subject \*\*\*diagnosed\*\*\* as having a disease assocd. with reduced CTL activity comprises administering a CTL activity-stimulating amt. of peptide T or an analog thereof. A method of treating a subject \*\*\*diagnosed\*\*\* as having a disease assocd. with reduced IFN-.gamma. activity comprises administering an IFN-.gamma. activity-stimulating amt. of peptide T or an analog thereof. A method of treating a subject \*\*\*diagnosed\*\*\* as having a disease assocd. with reduced IL-2 activity comprises administering an IL-2 activity-stimulating amt. of peptide T or an analog thereof.

TI Peptide T and analogs thereof for the stimulation of cytotoxic T lymphocyte (CTL) responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use

AB . . . in a subject, comprising administering a CTL activity-stimulating amt. of peptide T or an analog thereof. A method of increasing .gamma.- \*\*\*interferon\*\*\* (IFN-.gamma.) secretion in a subject comprises administering an IFN-.gamma. secretion-increasing amt. of peptide T or an analog thereof. A method of increasing \*\*\*interleukin\*\*\* 2 (IL-2) secretion in a subject comprises administering an IL-2 secretion-increasing amt. of peptide T or an analog thereof. A method of treating a subject \*\*\*diagnosed\*\*\* as having a disease assocd. with reduced CTL activity comprises administering a CTL activity-stimulating amt. of peptide T or an analog thereof. A method of treating a subject \*\*\*diagnosed\*\*\* as having a disease assocd. with reduced IFN-.gamma. activity comprises administering an IFN-.gamma. activity-stimulating amt. of peptide T or an analog thereof. A method of treating a subject \*\*\*diagnosed\*\*\* as having a disease assocd. with reduced IL-2 activity comprises administering an IL-2 activity-stimulating amt. of peptide T or an . . .

ST peptide T analog cytotoxic T lymphocyte response stimulation; \*\*\*interferon\*\*\* gamma secretion peptide T; \*\*\*interleukin\*\*\* 2 secretion peptide T

IT Lymphoma  
(B-cell; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Lymphoma  
(T-cell; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Tumor necrosis factors  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(TNF-.alpha.; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
(adenocarcinoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT AIDS (disease)  
(and AIDS-related lymphoma or sarcoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Infection  
(bacterial; peptide T and analogs for stimulation of cytotoxic T

lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Neoplasm  
 (blastoma; peptide T and analogs for stimulation of cytotoxic T  
 lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Urogenital system  
 (cancer; peptide T and analogs for stimulation of cytotoxic T  
 lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Esophagus, neoplasm  
 Head and Neck, neoplasm  
 Head and Neck, neoplasm  
 (carcinoma; peptide T and analogs for stimulation of cytotoxic T  
 lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Uterus, neoplasm  
 (cervix, carcinoma; peptide T and analogs for stimulation of cytotoxic  
 T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
 Uterus, neoplasm  
 (cervix; peptide T and analogs for stimulation of cytotoxic T  
 lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Intestine, neoplasm  
 (colon; peptide T and analogs for stimulation of cytotoxic T lymphocyte  
 responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and  
 \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Intestine, neoplasm  
 (colorectal; peptide T and analogs for stimulation of cytotoxic T  
 lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT T cell (lymphocyte)  
 (cytotoxic; peptide T and analogs for stimulation of cytotoxic T  
 lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
 (esophageal; peptide T and analogs for stimulation of cytotoxic T  
 lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Mycosis  
 (fungoides; peptide T and analogs for stimulation of cytotoxic T  
 lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Neuroglia, neoplasm  
 (glioblastoma; peptide T and analogs for stimulation of cytotoxic T  
 lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
 Carcinoma  
 (head and neck squamous cell carcinoma; peptide T and analogs for  
 stimulation of cytotoxic T lymphocyte responses and increasing  
 secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2,  
 and therapeutic use)

IT Carcinoma  
 Carcinoma

(head and neck; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Neoplasm  
(histiocytoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Hypoxia  
(hypoxic tumors; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Fungi  
Parasite  
(infection; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
(laryngeal squamous cell; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Neoplasm  
(metastasis; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Skin, neoplasm  
(mycosis fungoides; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Histiocyte  
(neoplasm, histiocytoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Nerve, neoplasm  
(neuroblastoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Lymphoma  
(non-Hodgkin's; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
(oral squamous cell; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Actinobacillus pleuropneumoniae  
Adenoma  
Alternaria alternata  
Anti-AIDS agents  
Antibacterial agents  
Antimalarials  
Antitumor agents  
Antiviral agents  
Aspergillus fumigatus

Bacillus anthracis  
Bladder, neoplasm  
Blastomyces dermatitidis  
Brain, neoplasm  
Brucella  
Brucella melitensis  
CD8-positive T cell  
Campylobacter  
Candida albicans  
Carcinoma  
Carcinoma  
Chlamydia pneumoniae  
Chlamydia trachomatis  
Chlamydophila psittaci  
Clostridium  
Clostridium tetani  
Coccidioides immitis  
Coronavirus  
Coxiella burnetii  
Cryptococcus neoformans  
Dengue virus  
Eastern equine encephalitis virus  
Ebola virus  
Ehrlichia  
Ehrlichia ruminantium  
Entamoeba histolytica  
Escherichia coli  
Fungicides  
Haemophilus  
Haemophilus ducreyi  
Haemophilus influenzae  
Hantavirus  
Hematopoietic neoplasm  
Hepatitis A virus  
Hepatitis B virus  
Hepatitis C virus  
Hepatitis E virus  
Hepatitis delta virus  
Histoplasma capsulatum  
Hodgkin's disease  
Human  
Human T-lymphotropic virus 1  
Human adenovirus  
Human coxsackievirus  
Human immunodeficiency virus  
Human immunodeficiency virus 1  
Human immunodeficiency virus 2  
Human papillomavirus  
Human poliovirus  
Immunostimulants  
Influenza A virus  
Influenza B virus  
Japanese encephalitis virus  
Kidney, neoplasm  
Lassa virus  
Legionella  
Legionella pneumophila

Leishmania  
Leishmania major  
Leukemia  
Listeria ivanovii  
Listeria monocytogenes  
Liver, neoplasm  
Lung, neoplasm  
Lymphoma  
Mammary gland, neoplasm  
Mannheimia haemolytica  
Marburg virus  
Measles virus  
Melanoma  
Multiple myeloma  
Mumps virus  
Murray Valley encephalitis virus  
Mycobacterium BCG  
Mycobacterium africanum  
Mycobacterium avium  
Mycobacterium avium \*\*\*paratuberculosis\*\*\*  
Mycobacterium bovis  
Mycobacterium intracellulare  
Mycobacterium kansasii  
Mycobacterium marinum  
Mycobacterium tuberculosis  
Mycobacterium ulcerans  
Myeloid leukemia  
Neisseria gonorrhoeae  
Neisseria meningitidis  
Neoplasm  
Nervous system, neoplasm  
Neuroglia, neoplasm  
Nocardia  
Nocardia asteroides  
Ovary, neoplasm  
Pancreas, neoplasm  
Paracoccidioides brasiliensis  
Parasiticides  
Pasteurella  
Pasteurella multocida  
Penicillium marneffeii  
Plasmodium (malarial genus)  
Plasmodium falciparum  
Plasmodium malariae  
Plasmodium vivax  
Pneumocystis carinii  
Polyomavirus  
Prostate gland, neoplasm  
Pseudomonas  
Pseudomonas aeruginosa  
Rabies virus  
Respiratory syncytial virus  
Rhinovirus  
Rickettsia  
Rift Valley fever virus  
Rotavirus A  
Rotavirus B

Rotavirus C  
 Rous sarcoma virus  
 Rubella virus  
 Salmonella  
 Salmonella typhi  
 Sarcoma  
 Schistosoma  
 Schistosoma mansoni  
 Shigella  
 Simian immunodeficiency virus  
 Sindbis virus  
 Skin, neoplasm  
 St. Louis encephalitis virus  
 Staphylococcus aureus  
 Staphylococcus epidermidis  
 Streptococcus agalactiae  
 Streptococcus pyogenes  
 Testis, neoplasm  
 Toxoplasma gondii  
 Trypanosoma brucei  
 Trypanosoma cruzi  
 Variola virus  
 Vesicular stomatitis virus  
 Vibrio cholerae  
 West Nile virus  
 Yellow fever virus  
 Yersinia  
 Yersinia enterocolitica  
 Yersinia pestis  
 (peptide T and analogs for stimulation of cytotoxic T lymphocyte  
 responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and  
 \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Cytokines  
 \*\*\*Interleukin\*\*\* 1  
 \*\*\*Interleukin\*\*\* 10  
 \*\*\*Interleukin\*\*\* 12  
 \*\*\*Interleukin\*\*\* 13  
 \*\*\*Interleukin\*\*\* 2  
 \*\*\*Interleukin\*\*\* 4  
 \*\*\*Interleukin\*\*\* 6  
 \*\*\*Interleukin\*\*\* 8  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (peptide T and analogs for stimulation of cytotoxic T lymphocyte  
 responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and  
 \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Peptides, biological studies  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
 (Biological study); USES (Uses)  
 (peptide T and analogs for stimulation of cytotoxic T lymphocyte  
 responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and  
 \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
 (pulmonary squamous cell; peptide T and analogs for stimulation of  
 cytotoxic T lymphocyte responses and increasing secretion of  
 \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and  
 therapeutic use)



IT Neoplasm  
(solid, carcinoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Head and Neck, neoplasm  
Head and Neck, neoplasm  
Larynx, neoplasm  
Lung, neoplasm  
Mouth, neoplasm  
(squamous cell carcinoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
(squamous cell; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Pharynx, neoplasm  
(throat squamous cell carcinoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Infection  
(viral; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT \*\*\*Interferons\*\*\*  
RL: BSU (Biological study, unclassified); BIOL (Biological study) (.gamma.; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT 106362-32-7D, C-terminal derivs.  
RL: PAC (Pharmacological activity); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT 106362-32-7, Peptide T 106362-32-7D, Peptide T, analogs  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT 107531-09-9 107531-11-3 107531-12-4 107531-14-6 118936-25-7  
118936-26-8 118936-27-9 118936-30-4 118936-31-5 118936-32-6  
118957-86-1 119386-95-7  
RL: PRP (Properties)  
(unclaimed sequence; peptide T and analogs thereof for the stimulation of cytotoxic T lymphocyte (CTL) responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

DN PubMed ID: 12208110  
 TI Localisation of CD25+ cells and MHCII+ cells in lymph nodes draining  
 Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* vaccination  
 granuloma and the presence of a systemic immune response.  
 AU Valheim M; Hasvold H J; Storset A K; Larsen H J S; Press C McL  
 CS Department of Morphology, Genetics and Aquatic Biology, Norwegian School  
 of Veterinary Science, P.O. Box 8146 Dep. N-0033 Oslo, Norway..  
 mette.valheim@vetinst.no  
 SO Research in veterinary science, (2002 Aug) Vol. 73, No. 1, pp. 77-85.  
 Journal code: 0401300. ISSN: 0034-5288.  
 CY England: United Kingdom  
 DT (CLINICAL TRIAL)  
 Journal; Article; (JOURNAL ARTICLE)  
 LA English  
 FS Priority Journals  
 EM 200305  
 ED Entered STN: 6 Sep 2002  
 Last Updated on STN: 21 May 2003  
 Entered Medline: 20 May 2003  
 AB Vaccination of goat kids against \*\*\*paratuberculosis\*\*\* protects  
 against lesions and clinical disease. The systemic cellular response was  
 studied in goat kids 3-9 weeks after vaccination. Peripheral blood cells  
 showed increased \*\*\*interferon\*\*\* -gamma production and expression of  
 \*\*\*interleukin\*\*\* -2 receptor (CD25) after stimulation with  
 Mycobacterium  
 avium subsp. \*\*\*paratuberculosis\*\*\* antigens. The lymph node draining  
 the vaccination granuloma was studied three weeks after vaccination in a  
 parallel group of goat kids. In deep cortex, MHCII+ cells were observed  
 surrounded by CD4+ T-cells, while follicular hypertrophy and hyperplasia  
 were prominent in the subcapsular region and along connective tissue  
 trabecula. Comparison of the local and systemic immune responses revealed  
 an inverse relationship between CD25+ T-cells in the lymph node deep  
 cortex and cells in peripheral blood that up-regulate CD25 upon in vitro  
 stimulation, suggesting that activated and regulatory T-cells in the local  
 lymph node influence the level of circulating antigen-specific T-cells  
 following vaccination against \*\*\*paratuberculosis\*\*\* in goats.  
 TI Localisation of CD25+ cells and MHCII+ cells in lymph nodes draining  
 Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* vaccination  
 granuloma and the presence of a systemic immune response.  
 AB Vaccination of goat kids against \*\*\*paratuberculosis\*\*\* protects  
 against lesions and clinical disease. The systemic cellular response was  
 studied in goat kids 3-9 weeks after vaccination. Peripheral blood cells  
 showed increased \*\*\*interferon\*\*\* -gamma production and expression of  
 \*\*\*interleukin\*\*\* -2 receptor (CD25) after stimulation with  
 Mycobacterium  
 avium subsp. \*\*\*paratuberculosis\*\*\* antigens. The lymph node draining  
 the vaccination granuloma was studied three weeks after vaccination in a  
 parallel group of goat. . . that activated and regulatory T-cells in  
 the local lymph node influence the level of circulating antigen-specific  
 T-cells following vaccination against \*\*\*paratuberculosis\*\*\* in goats.  
 CT Check Tags: Male  
 Animals  
 Antigens, Bacterial: IM, immunology  
 \*Bacterial Vaccines: IM, immunology  
 \*\*\* Enzyme-Linked Immunosorbent Assay\*\*\*  
 Gene Expression Regulation  
 Goat Diseases: IM, immunology

Goat Diseases: MI, microbiology  
 Goat Diseases: PA, pathology  
 Goat Diseases:. . . IM, immunology  
 Goats: MI, microbiology  
 \*Granuloma: IM, immunology  
 Granuloma: MI, microbiology  
 Granuloma: PA, pathology  
 \*Histocompatibility Antigens Class II: IM, immunology  
 \*\*\* Interferon Type II: IM, immunology\*\*\*  
 \*\*\* Interferon Type II: ME, metabolism\*\*\*  
 Lymph Nodes: CY, cytology  
 \*Lymph Nodes: IM, immunology  
 Lymph Nodes: MI, microbiology  
 \*\*\*\*Mycobacterium avium subsp. paratuberculosis: IM, immunology\*\*\*  
 \*\*\* Mycobacterium avium subsp. paratuberculosis: IP, isolation &\*\*\*  
 \*\*\* purification\*\*\*  
 \*\*\* Paratuberculosis: IM, immunology\*\*\*  
 \*\*\* Paratuberculosis: MI, microbiology\*\*\*  
 \*\*\* Paratuberculosis: PA, pathology\*\*\*  
 \*\*\* Paratuberculosis: PC, prevention & control\*\*\*  
 \*\*\* Receptors, Interleukin-2: AN, analysis\*\*\*  
 \*\*\*\*Receptors, Interleukin-2: IM, immunology\*\*\*  
 \*T-Lymphocytes: IM, immunology  
 RN \*\*\*82115-62-6 (Interferon Type II)\*\*\*  
 CN 0 (Antigens, Bacterial); 0 (Bacterial Vaccines); 0 (Histocompatibility  
 Antigens Class II); 0 (Receptors, \*\*\*Interleukin\*\*\* -2)

L10 ANSWER 13 OF 19 MEDLINE on STN  
 AN 2002648148 MEDLINE <<LOGINID::20080325>>  
 DN PubMed ID: 12406650  
 TI Type 1 and type 2 responses in regulation of Ig isotype expression in  
 cattle.  
 AU Estes D Mark; Brown Wendy C  
 CS Program for the Prevention of Animal Infectious Diseases, Department of  
 Veterinary Pathobiology, University of Missouri, Columbia, MO 65211, USA..  
 estesd@missouri.edu  
 SO Veterinary immunology and immunopathology, (2002 Nov) Vol. 90, No. 1-2,  
 pp. 1-10. Ref: 90  
 Journal code: 8002006. ISSN: 0165-2427.  
 CY Netherlands  
 DT Journal; Article; (JOURNAL ARTICLE)  
 General Review; (REVIEW)  
 LA English  
 FS Priority Journals  
 EM 200305  
 ED Entered STN: 31 Oct 2002  
 Last Updated on STN: 3 May 2003  
 Entered Medline: 2 May 2003  
 AB Regulation of humoral immune responses is multifactorial involving  
 appropriate activation, costimulation and the presence of specific soluble  
 factors. Polarized type 1 or type 2 humoral responses in the laboratory  
 mouse have been linked to expression of specific cytokines and thus can be  
 used to provide insight into the type of response generated by infection.  
 For example, IFN-gamma has been linked to IgG2a and IgG3 production, IL-4  
 to IgG1 and IgE production and TGF-beta to IgA production. Unlike the  
 laboratory mouse, generally housed under defined conditions, highly skewed  
 isotype expression patterns generally occur in cattle in chronic

infections. A few examples of polarized responses have been noted in chronic experimental or naturally occurring infections including F. hepatica, M. \*\*\*paratuberculosis\*\*\*, C. parvum and B. abortus. In vitro studies using purified bovine B cells and various forms of costimulation and cytokines have demonstrated that isotype responses can be polarized under certain experimental conditions in vitro. That is, IgG1 expression is positively regulated by IL-4 and IgG2 expression is positively regulated by IFN-gamma. Other as yet unidentified factors may play pivotal roles in regulating humoral immune responses in large ruminant species in vivo. This possibility is best exemplified by recent studies using DNA vaccines in cattle that have been demonstrated in the mouse to be generally polarizing to a type 1 response. Surprisingly, studies in cattle using plasmid DNA as vaccination material show an almost exclusive IgG1 response. Based on a number of studies using T cell clones and various biological \*\*\*assays\*\*\*, it is clear that the classical roles of many cytokines in the laboratory mouse do not extrapolate entirely or at all to cattle. Thus, the design of adjuvants and immune modulators should be based on studies done in cattle or using bovine cells. Based on studies to date, several "holes" in the cytokine repertoire exist and these roles may be assumed by unique factors or activities of other known cytokines.

AB . . . A few examples of polarized responses have been noted in chronic experimental or naturally occurring infections including F. hepatica, M. \*\*\*paratuberculosis\*\*\*, C. parvum and B. abortus. In vitro studies using purified bovine B cells and various forms of costimulation and cytokines. . . material show an almost exclusive IgG1 response. Based on a number of studies using T cell clones and various biological \*\*\*assays\*\*\*, it is clear that the classical roles of many cytokines in the laboratory mouse do not extrapolate entirely or at. . .

CT Animals

\*Cattle: IM, immunology

\*Gene Expression Regulation

\*Immunoglobulin Isotypes: GE, genetics

\*\*\* Interferon Type II: IM, immunology\*\*\*

\*\*\* Interleukins: IM, immunology\*\*\*

\*Th1 Cells: IM, immunology

\*Th2 Cells: IM, immunology

Transforming Growth Factor beta: IM, immunology

RN \*\*\*82115-62-6 (Interferon Type II)\*\*\*

CN 0 (Immunoglobulin Isotypes); 0 ( \*\*\*Interleukins\*\*\* ); 0 (Transforming Growth Factor beta)

L10 ANSWER 14 OF 19 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN

AN 2001:570180 BIOSIS <<LOGINID::20080325>>

DN PREV200100570180

TI Results of multiple \*\*\*diagnostic\*\*\* tests for Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* in patients with inflammatory bowel disease and in controls.

AU Collins, Michael T. [Reprint author]; Lisby, Gorm; Moser, Claus; Chicks, Debra; Christensen, Steen; Reichelderfer, Mark; Hoiby, Niels; Harms, Bruce A.; Thomsen, Ole O.; Skibsted, Ulrik; Binder, Vibeke

CS Department of Pathobiological Sciences, School of Veterinary Medicine, 2015 Linden Dr. West, Madison, WI, 53706-1102, USA  
mcollin5@facstaff.wisc.edu

SO Journal of Clinical Microbiology, (December, 2001) Vol. 38, No. 12, pp. 4373-4381. print.

CODEN: JCMIDW. ISSN: 0095-1137.

DT Article

LA English

ED Entered STN: 12 Dec 2001

Last Updated on STN: 25 Feb 2002

AB Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* has been incriminated as a cause of Crohn's disease (CD); however, studies to date have been relatively small and generally only used a single \*\*\*diagnostic\*\*\* \*\*\*assay\*\*\*. The objective of the study was to reexamine the association of M. avium subsp. \*\*\*paratuberculosis\*\*\* and CD using multiple \*\*\*diagnostic\*\*\* tests. Five methods were used to detect M. avium subsp. \*\*\*paratuberculosis\*\*\* infections in 439 inflammatory bowel disease (IBD) patients and 324 control subjects in the United States and Denmark. Most \*\*\*assays\*\*\* were adaptations of \*\*\*diagnostic\*\*\* tests for this infection performed routinely on animals. PCR for IS900, a genetic element unique to M. avium subsp. \*\*\*paratuberculosis\*\*\*, was positive significantly more often on resected bowel and lymph node tissues from CD patients (19.0%) and ulcerative colitis (UC) patients (26.2%) than from controls (6.3%) ( $P < 0.05$ ). Positive IS900 PCR results occurred more often in U.S. than in Danish IBD patients, 32.0 versus 13.3% ( $P = 0.025$ ). The majority of Danish patients were bacillus Calmette-Guerin (Mycobacterium bovis BCG) vaccinated (CD, 77.5%; UC, 86.6%; controls, 83.0%) whereas none of the U.S. patients with IBD and only 2% of U.S. controls were vaccinated. Among Danish IBD patients, positive PCR findings were four times more common among subjects who were not BCG vaccinated (33.3%) than among BCG vaccinates (8.8%,  $P = 0.02$ ). Culture of the same tissues tested by PCR using modified BACTEC 12B medium failed to grow M. avium subsp. \*\*\*paratuberculosis\*\*\* from patients or controls. U.S. CD patients

had

the highest serological evidence (enzyme-linked immunosorbent \*\*\*assay\*\*\* (ELISA) for serum antibodies) of M. avium subsp. \*\*\*paratuberculosis\*\*\* infection (20.7% of patients positive) which was higher than for all UC patients studied (6.1%) or healthy controls (3.8%,  $P < 0.005$ ). Among Danish patients alone, however, no significant differences in rates of ELISA-positive results among CD, UC, or control patients were found. For 181 study subjects, both IS900 PCR and ELISA were performed. Although 11 were ELISA positive and 36 were PCR positive, in no instance was a patient positive by both tests, suggesting that these states are mutually exclusive. Evaluation of cytokine-mediated immune responses of IBD patients was complicated by the influence of immunosuppressive therapy given most IBD patients. Gamma \*\*\*interferon\*\*\* (IFN-gamma) release by peripheral blood leukocytes after M. avium purified protein derivative PPD antigen stimulation showed significantly lower responses in CD patients than in UC patients or controls in both U.S. (by ex vivo \*\*\*assay\*\*\* ) and Danish (by in vitro \*\*\*assay\*\*\* ) populations ( $P < 0.05$ ). \*\*\*Interleukin\*\*\* -5 responses were not different among CD, UC, or control groups. Collectively, the PCR, ELISA, and IFN-gamma tests for M. avium subsp.

BCG

\*\*\*paratuberculosis\*\*\* together with the unexpected observation that vaccination influenced M. avium subsp. \*\*\*paratuberculosis\*\*\* detection, lead us to conclude that M. avium subsp.

\*\*\*paratuberculosis\*\*\*, or some similarly fastidious mycobacterial species, infects at least a subset of IBD patients. Whether the infection is primary (causal) or secondary, it may contribute to the etiopathogenesis of IBD.

TI Results of multiple \*\*\*diagnostic\*\*\* tests for Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* in patients with inflammatory bowel disease and in controls.

AB Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* has been incriminated as a cause of Crohn's disease (CD); however, studies to date have been relatively small and generally only used a single \*\*\*diagnostic\*\*\* \*\*\*assay\*\*\*. The objective of the study was to reexamine the association of M. avium subsp. \*\*\*paratuberculosis\*\*\* and CD using multiple \*\*\*diagnostic\*\*\* tests. Five methods were used to detect M. avium subsp. \*\*\*paratuberculosis\*\*\* infections in 439 inflammatory bowel disease (IBD) patients and 324 control subjects in the United States and Denmark. Most \*\*\*assays\*\*\* were adaptations of \*\*\*diagnostic\*\*\* tests for this infection performed routinely on animals. PCR for IS900, a genetic element unique to M. avium subsp. \*\*\*paratuberculosis\*\*\*, was positive significantly more often on resected bowel and lymph node tissues from CD patients (19.0%) and ulcerative colitis (UC). . . . P=0.02). Culture of the same tissues tested by PCR using modified BACTEC 12B medium failed to grow M. avium subsp. \*\*\*paratuberculosis\*\*\* from patients or controls. U.S. CD patients had the highest serological evidence (enzyme-linked immunosorbent \*\*\*assay\*\*\* (ELISA) for serum antibodies) of M. avium subsp. \*\*\*paratuberculosis\*\*\* infection (20.7% of patients positive) which was higher than for all UC patients studied (6.1%) or healthy controls (3.8%, P<0.005).. . . of cytokine-mediated immune responses of IBD patients was complicated by the influence of immunosuppressive therapy given most IBD patients. Gamma \*\*\*interferon\*\*\* (IFN-gamma) release by peripheral blood leukocytes after M. avium purified protein derivative PPD antigen stimulation showed significantly lower responses in CD patients than in UC patients or controls in both U.S. (by ex vivo \*\*\*assay\*\*\* ) and Danish (by in vitro \*\*\*assay\*\*\* ) populations (P<0.05). \*\*\*Interleukin\*\*\* -5 responses were not different among CD, UC, or control groups. Collectively, the PCR, ELISA, and IFN-gamma tests for M. avium subsp. \*\*\*paratuberculosis\*\*\* together with the unexpected observation that BCG vaccination influenced M. avium subsp. \*\*\*paratuberculosis\*\*\* detection, lead us to conclude that M. avium subsp. \*\*\*paratuberculosis\*\*\*, or some similarly fastidious mycobacterial species, infects at least a subset of IBD patients. Whether the infection is primary (causal). . .

IT . . .  
     system disease  
     Crohn Disease (MeSH)

IT Diseases  
     inflammatory bowel disease: digestive system disease  
     Inflammatory Bowel Diseases (MeSH)

IT Chemicals & Biochemicals  
     IFN-gamma [ \*\*\*interferon\*\*\* -gamma]; PPD antigen

IT Methods & Equipment  
     ELISA: analytical method, labeling; PCR [polymerase chain reaction]: DNA amplification, \*\*\*diagnostic\*\*\* method, in situ recombinant gene expression detection, sequencing techniques

ORGN . . .  
     Vertebrates

ORGN Classifier  
     Mycobacteriaceae 08881  
     Super Taxa  
     Mycobacteria; Actinomycetes and Related Organisms; Eubacteria; Bacteria; Microorganisms

Organism Name

Mycobacterium avium ssp. \*\*\*paratuberculosis\*\*\* : pathogen

Mycobacterium bovis: pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

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STN DUPLICATE 5

AN 2001:410237 BIOSIS <<LOGINID::20080325>>

DN PREV200100410237

TI Subclinical \*\*\*paratuberculosis\*\*\* in goats following experimental  
infection: An immunological and microbiological study.

AU Storset, A. K. [Reprint author]; Hasvold, H. J.; Valheim, M.; Brun-Hansen,  
H.; Berntsen, G.; Whist, S. K.; Djonne, B.; Press, C. McL.; Holstad, G.;  
Larsen, H. J. S.

CS Department of Pharmacology, Microbiology and Food Hygiene, Norwegian  
School of Veterinary Science, N-0033, Oslo, Norway  
anne.storset@veths.no

SO Veterinary Immunology and Immunopathology, (10 August, 2001) Vol. 80, No.  
3-4, pp. 271-287. print.  
CODEN: VIIMDS. ISSN: 0165-2427.

DT Article

LA English

ED Entered STN: 29 Aug 2001

Last Updated on STN: 22 Feb 2002

AB An experimental oral infection of goats with a caprine isolate of  
Mycobacterium a. subsp. \*\*\*paratuberculosis\*\*\* was used to investigate  
immunological and bacteriological events during the subclinical phase of  
infection. Seven goats at 5-8 weeks of age were given a bacterial  
suspension in milk-replacement three times weekly for 9 weeks. Six  
animals were kept as controls. Cellular recall responses against M. a.  
\*\*\*paratuberculosis\*\*\* were analysed by means of a lymphocyte  
proliferation test, an IFN-gamma \*\*\*assay\*\*\* and an IL-2 receptor  
\*\*\*assay\*\*\*. All inoculated animals had detectable CMI responses from

9

weeks post-inoculation and through the 2 years of study, although the  
responses were highest during the first year. Antibodies against M. a.

\*\*\*paratuberculosis\*\*\* could be detected from weeks 15-20 in four of  
the

seven animals, and one additional animal became antibody positive at week  
35, while two inoculated animals did not produce significant antibody  
titres during the experiment. At about 1-year post-inoculation, two  
animals became faecal shedders, while two others started to excrete  
bacteria into faeces about 2 years post-inoculation. The appearance of M.  
a. \*\*\*paratuberculosis\*\*\* in faeces was not associated with a decline  
in cellular responses as far as could be assessed using the current  
methods for measuring CMI. Pathological lesions due to M. a.

\*\*\*paratuberculosis\*\*\* infection and presence of bacteria were recorded  
in the intestine and/or mesenteric lymph nodes of five animals while lymph  
node changes suggestive of \*\*\*paratuberculosis\*\*\* were observed in one  
animal. Only the two animals with no signs of an active infection at  
necropsy showed a considerable decline in the cellular parameters during  
the last year of the study, particularly in the IFN-gamma \*\*\*assay\*\*\*.  
The two animals with the highest levels of M. a. \*\*\*paratuberculosis\*\*\*  
responsive CD8+ lymphocytes in the circulation about 1-year  
post-inoculation had no detectable lesions in the distal ileum and colon  
at necropsy, while high numbers of gammadelta T-cells responsive to M. a.

\*\*\*paratuberculosis\*\*\* in the circulation were associated with disseminated lesions in the distal ileum and colon.

TI Subclinical \*\*\*paratuberculosis\*\*\* in goats following experimental infection: An immunological and microbiological study.

AB An experimental oral infection of goats with a caprine isolate of *Mycobacterium a. subsp.* \*\*\*paratuberculosis\*\*\* was used to investigate immunological and bacteriological events during the subclinical phase of infection. Seven goats at 5-8 weeks of . . . in milk-replacement three times weekly for 9 weeks. Six animals were kept as controls. Cellular recall responses against *M. a.* \*\*\*paratuberculosis\*\*\* were analysed by means of a lymphocyte proliferation test, an IFN-gamma \*\*\*assay\*\*\* and an IL-2 receptor \*\*\*assay\*\*\*. All inoculated animals had detectable CMI responses from 9 weeks post-inoculation and through the 2 years of study, although the responses were highest during the first year. Antibodies against *M. a.* \*\*\*paratuberculosis\*\*\* could be detected from weeks 15-20 in four of the seven animals, and one additional animal became antibody positive at. . . faecal shedders, while two others started to excrete bacteria into faeces about 2 years post-inoculation. The appearance of *M. a.* \*\*\*paratuberculosis\*\*\* in faeces was not associated with a decline in cellular responses as far as could be assessed using the current methods for measuring CMI. Pathological lesions due to *M. a.* \*\*\*paratuberculosis\*\*\* infection and presence of bacteria were recorded in the intestine and/or mesenteric lymph nodes of five animals while lymph node changes suggestive of \*\*\*paratuberculosis\*\*\* were observed in one animal. Only the two animals with no signs of an active infection at necropsy showed a considerable decline in the cellular parameters during the last year of the study, particularly in the IFN-gamma \*\*\*assay\*\*\*. The two animals with the highest levels of *M. a.* \*\*\*paratuberculosis\*\*\* responsive CD8+ lymphocytes in the circulation about 1-year post-inoculation had no detectable lesions in the distal ileum and colon at necropsy, while high numbers of gammadelta T-cells responsive to *M. a.* \*\*\*paratuberculosis\*\*\* in the circulation were associated with disseminated lesions in the distal ileum and colon.

IT . . .  
digestive system; lymphocyte: blood and lymphatics, immune system;  
mesenteric lymph node: blood and lymphatics, digestive system, immune system

IT Diseases  
\*\*\*paratuberculosis\*\*\* : bacterial disease, subclinical  
\*\*\*Paratuberculosis\*\*\* (MeSH)

IT Chemicals & Biochemicals  
IFN-gamma [ \*\*\*interferon\*\*\* -gamma]; IL-2 receptor [ \*\*\*interleukin\*\*\* -2 receptor]

IT Methods & Equipment  
IFN-gamma \*\*\*assay\*\*\* : analytical method; \*\*\*interleukin\*\*\* -2 receptor \*\*\*assay\*\*\* : analytical method; lymphocyte proliferation test: analytical method; necropsy: analytical method

IT Miscellaneous Descriptors  
cellular immunity

ORGN . . .  
Mammals, Vertebrates

ORGN Classifier  
Mycobacteriaceae 08881  
Super Taxa  
Mycobacteria; Actinomycetes and Related Organisms; Eubacteria;  
Bacteria; Microorganisms



Organism Name

Mycobacterium avium \*\*\*paratuberculosis\*\*\* : pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

L10 ANSWER 16 OF 19 LIFESCI COPYRIGHT 2008 CSA on STN DUPLICATE 6  
AN 2001:53334 LIFESCI <<LOGINID::20080325>>  
TI Results of Multiple \*\*\*Diagnostic\*\*\* Tests for Mycobacterium avium  
subsp. \*\*\*paratuberculosis\*\*\* in Patients with Inflammatory Bowel  
Disease and in Controls  
AU Collins\*, M.T.; Lisby, G.; Moser, C.; Chicks, D.; Christensen, S.;  
Reichelderfer, M.; Hoeiby, N.; Harms, B.A.; Thomsen, O.O.; Skibsted, U.;  
Binder, V.  
CS Department of Pathobiological Sciences, School of Veterinary Medicine,  
2015 Linden Dr. West, Madison, WI 53706-1102; E-mail:  
mcollin5@facstaff.wisc.edu  
SO Journal of Clinical Microbiology [J. Clin. Microbiol.], (20001200) vol.  
38, no. 12, pp. 4373-4381.  
ISSN: 0095-1137.  
DT Journal  
FS J; A  
LA English  
SL English  
AB Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* has been  
incriminated as a cause of Crohn's disease (CD); however, studies to date  
have been relatively small and generally only used a single  
\*\*\*diagnostic\*\*\* \*\*\*assay\*\*\*. The objective of the study was to  
reexamine the association of M. avium subsp. \*\*\*paratuberculosis\*\*\*  
and CD using multiple \*\*\*diagnostic\*\*\* tests. Five methods were used  
to detect Mavium subsp. \*\*\*paratuberculosis\*\*\* infections in 439  
inflammatory bowel disease (IBD) patients and 324 control subjects in the  
United States and Denmark. Most \*\*\*assays\*\*\* were adaptations of  
\*\*\*diagnostic\*\*\* tests for this infection performed routinely on  
animals. PCR for IS900, a genetic element unique to M. avium subsp.  
\*\*\*paratuberculosis\*\*\*, was positive significantly more often on  
resected bowel and lymph node tissues from CD patients (19.0%) and  
ulcerative colitis (UC) patients (26.2%) than from controls (6.3%) ( $P < 0.05$ ). Positive IS900 PCR results occurred more often in U.S. than in  
Danish IBD patients, 32.0 versus 13.3% ( $P = 0.025$ ). The majority of Danish  
patients were bacillus Calmette-Guerin (Mycobacterium bovis BCG)  
vaccinated (CD, 77.5%; UC, 86.6%; controls, 83.0%) whereas none of the  
U.S. patients with IBD and only 2% of U.S. controls were vaccinated. Among  
Danish IBD patients, positive PCR findings were four times more common  
among subjects who were not BCG vaccinated (33.3%) than among BCG  
vaccinates (8.8%,  $P = 0.02$ ). Culture of the same tissues tested by PCR  
using modified BACTEC 12B medium failed to grow M. avium subsp.  
\*\*\*paratuberculosis\*\*\* from patients or controls. U.S. CD patients had  
the highest serological evidence (enzyme-linked immunosorbent  
\*\*\*assay\*\*\* [ELISA] for serum antibodies) of M. avium subsp.  
\*\*\*paratuberculosis\*\*\* infection (20.7% of patients positive) which was  
higher than for all UC patients studied (6.1%) or healthy controls (3.8%,  
 $P < 0.005$ ). Among Danish patients alone, however, no significant  
differences in rates of ELISA-positive results among CD, UC, or control  
patients were found. For 181 study subjects, both IS900 PCR and ELISA were  
performed. Although 11 were ELISA positive and 36 were PCR positive, in no  
instance was a patient positive by both tests, suggesting that these  
states are mutually exclusive. Evaluation of cytokine-mediated immune

responses of IBD patients was complicated by the influence of immunosuppressive therapy given most IBD patients. Gamma

\*\*\*interferon\*\*\* (IFN- gamma ) release by peripheral blood leukocytes after M. avium purified protein derivative PPD antigen stimulation showed significantly lower responses in CD patients than in UC patients or controls in both U.S. (by ex vivo \*\*\*assay\*\*\* ) and Danish (by in vitro \*\*\*assay\*\*\* ) populations (P < 0.05). \*\*\*Interleukin\*\*\* -5 responses were not different among CD, UC, or control groups. Collectively, the PCR, ELISA, and IFN- gamma tests for M. avium subsp. \*\*\*paratuberculosis\*\*\* together with the unexpected observation that BCG vaccination influenced M. avium subsp. \*\*\*paratuberculosis\*\*\* detection, lead us to conclude that M. avium subsp. \*\*\*paratuberculosis\*\*\* , or some similarly fastidious mycobacterial species, infects at least a subset of IBD patients. Whether the infection is primary (causal) or secondary, it may contribute to the etiopathogenesis of IBD.

TI Results of Multiple \*\*\*Diagnostic\*\*\* Tests for Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* in Patients with Inflammatory Bowel Disease and in Controls

AB Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* has been incriminated as a cause of Crohn's disease (CD); however, studies to date have been relatively small and generally only used a single

\*\*\*diagnostic\*\*\* \*\*\*assay\*\*\* . The objective of the study was to reexamine the association of M. avium subsp. \*\*\*paratuberculosis\*\*\* and CD using multiple \*\*\*diagnostic\*\*\* tests. Five methods were used to detect M. avium subsp. \*\*\*paratuberculosis\*\*\* infections in 439 inflammatory bowel disease (IBD) patients and 324 control subjects in the United States and Denmark. Most \*\*\*assays\*\*\* were adaptations of \*\*\*diagnostic\*\*\* tests for this infection performed routinely on animals. PCR for IS900, a genetic element unique to M. avium subsp.

\*\*\*paratuberculosis\*\*\* , was positive significantly more often on resected bowel and lymph node tissues from CD patients (19.0%) and ulcerative colitis (UC). . . . 0.02). Culture of the same tissues tested by PCR using modified BACTEC 12B medium failed to grow M. avium subsp.

\*\*\*paratuberculosis\*\*\* from patients or controls. U.S. CD patients had the highest serological evidence (enzyme-linked immunosorbent \*\*\*assay\*\*\* [ELISA] for serum antibodies) of M. avium subsp.

\*\*\*paratuberculosis\*\*\* infection (20.7% of patients positive) which was higher than for all UC patients studied (6.1%) or healthy controls (3.8%, P. . . . of cytokine-mediated immune responses of IBD patients was complicated by the influence of immunosuppressive therapy given most IBD patients. Gamma \*\*\*interferon\*\*\* (IFN- gamma ) release by peripheral blood leukocytes after M. avium purified protein derivative PPD antigen stimulation showed significantly lower responses in CD patients than in UC patients or controls in both U.S. (by ex vivo \*\*\*assay\*\*\* ) and Danish (by in vitro \*\*\*assay\*\*\* ) populations (P < 0.05). \*\*\*Interleukin\*\*\* -5 responses were not different among CD, UC, or control groups. Collectively, the PCR, ELISA, and IFN- gamma tests for M. avium subsp.

\*\*\*paratuberculosis\*\*\* together with the unexpected observation that

BCG

vaccination influenced M. avium subsp. \*\*\*paratuberculosis\*\*\* detection, lead us to conclude that M. avium subsp.

\*\*\*paratuberculosis\*\*\* , or some similarly fastidious mycobacterial species, infects at least a subset of IBD patients. Whether the infection is primary (causal). . . .

UT Bactec test; BCG; Enzyme-linked immunosorbent \*\*\*assay\*\*\* ; Polymerase chain reaction; \*\*\*Diagnostic\*\*\* agents; Inflammatory bowel diseases; Crohn's disease; USA; Denmark; Mycobacterium avium

\*\*\*paratuberculosis\*\*\* ; tests; man

L10 ANSWER 17 OF 19 MEDLINE on STN  
AN 2001023445 MEDLINE <<LOGINID::20080325>>  
DN PubMed ID: 10895895  
TI Cytokine secretion by peripheral blood mononuclear cells from cows  
infected with Mycobacterium \*\*\*paratuberculosis\*\*\* .  
AU Stabel J R  
CS Bacterial Diseases of Livestock Research Unit, USDA-Agricultural Research  
Services, National Animal Disease Center, Ames, IA 50010, USA.  
SO American journal of veterinary research, (2000 Jul) Vol. 61, No. 7, pp.  
754-60.  
Journal code: 0375011. ISSN: 0002-9645.  
CY United States  
DT Journal; Article; (JOURNAL ARTICLE)  
LA English  
FS Priority Journals  
EM 200011  
ED Entered STN: 22 Mar 2001  
Last Updated on STN: 22 Mar 2001  
Entered Medline: 3 Nov 2000  
AB OBJECTIVE: To compare cytokine secretion patterns of peripheral blood  
mononuclear cells (PBMC) from healthy cows and cows subclinically and  
clinically infected with Mycobacterium \*\*\*paratuberculosis\*\*\* .  
ANIMALS: 5 noninfected cows, 6 cows with subclinical  
\*\*\*paratuberculosis\*\*\* , and 4 cows with clinical  
\*\*\*paratuberculosis\*\*\* . PROCEDURE: PBMC were isolated, and  
concentrations or activities of secreted \*\*\*interleukin\*\*\* (IL)-1,  
IL-2, IL-6, tumor necrosis factor (TNF), and \*\*\*interferon\*\*\* -gamma  
(IFN-gamma) were measured after in vitro stimulation of cells with  
concanavalin A (ConA), lipopolysaccharide (LPS), or a whole-cell sonicate  
of M \*\*\*paratuberculosis\*\*\* (MpS). Proliferative responses of PBMC  
were also determined after stimulation with ConA, phytohemagglutinin,  
pokeweed mitogen (PWM), or MpS. RESULTS: After stimulation with ConA,  
cells from subclinically infected cows secreted significantly more, and  
cells from clinically infected cows secreted significantly less,  
IFN-gamma, compared with cells from control cows. Cells from cows with  
subclinical \*\*\*paratuberculosis\*\*\* produced significantly more TNF and  
IFN-gamma in response to MpS than cells from the other 2 groups.  
Stimulation of PBMC from subclinically infected cows with ConA or MpS  
resulted in significantly higher proliferative responses, compared with  
cells from control and clinically infected cows. In contrast, clinically  
infected cows had significantly higher proliferative responses to PWM than  
cells from the other 2 groups. CONCLUSIONS AND CLINICAL RELEVANCE: A  
decrease in T-cell responses to mitogens or MpS was observed in cows  
clinically infected with M \*\*\*paratuberculosis\*\*\* , compared with  
subclinically infected cows, suggesting that activated T cells may delay  
the progression of \*\*\*paratuberculosis\*\*\* .  
TI Cytokine secretion by peripheral blood mononuclear cells from cows  
infected with Mycobacterium \*\*\*paratuberculosis\*\*\* .  
AB . . . cytokine secretion patterns of peripheral blood mononuclear cells  
(PBMC) from healthy cows and cows subclinically and clinically infected  
with Mycobacterium \*\*\*paratuberculosis\*\*\* . ANIMALS: 5 noninfected  
cows, 6 cows with subclinical \*\*\*paratuberculosis\*\*\* , and 4 cows with  
clinical \*\*\*paratuberculosis\*\*\* . PROCEDURE: PBMC were isolated, and  
concentrations or activities of secreted \*\*\*interleukin\*\*\* (IL)-1,  
IL-2, IL-6, tumor necrosis factor (TNF), and \*\*\*interferon\*\*\* -gamma

(IFN-gamma) were measured after in vitro stimulation of cells with concanavalin A (ConA), lipopolysaccharide (LPS), or a whole-cell sonicate of M \*\*\*paratuberculosis\*\*\* (MpS). Proliferative responses of PBMC were also determined after stimulation with ConA, phytohemagglutinin, pokeweed mitogen (PWM), or MpS. RESULTS: After. . . cells from clinically infected cows secreted significantly less, IFN-gamma, compared with cells from control cows. Cells from cows with subclinical \*\*\*paratuberculosis\*\*\* produced significantly more TNF and IFN-gamma in response to MpS than cells from the other 2 groups. Stimulation of PBMC. . . AND CLINICAL RELEVANCE: A decrease in T-cell responses to mitogens or MpS was observed in cows clinically infected with M \*\*\*paratuberculosis\*\*\*, compared with subclinically infected cows, suggesting that activated T cells may delay the progression of \*\*\*paratuberculosis\*\*\*.

CT . . .

\*Cattle Diseases: IM, immunology

Cattle Diseases: MI, microbiology

Cell Division

Concanavalin A: IM, immunology

Cytokines: AN, analysis

\*Cytokines: SE, secretion

\*\*\* Enzyme-Linked Immunosorbent Assay: VE, veterinary\*\*\*

\*\*\* Interferon Type II: AN, analysis\*\*\*

\*\*\* Interleukin-1: AN, analysis\*\*\*

\*\*\* Interleukin-2: AN, analysis\*\*\*

Leukocytes, Mononuclear: IM, immunology

\*Leukocytes, Mononuclear: SE, secretion

Lipopolysaccharides: IM, immunology

Lymphocyte Activation

\*\*\*\*Mycobacterium avium subsp. paratuberculosis: IM, immunology\*\*\*

\*\*\* Mycobacterium avium subsp. paratuberculosis: PY, pathogenicity\*\*\*

\*\*\* Paratuberculosis: BL, blood\*\*\*

\*\*\*\*Paratuberculosis: IM, immunology\*\*\*

\*\*\* Paratuberculosis: MI, microbiology\*\*\*

Phytohemagglutinins: IM, immunology

Pokeweed Mitogens: IM, immunology

Scintillation Counting: VE, veterinary

RN 11028-71-0 (Concanavalin A); \*\*\*82115-62-6 (Interferon Type II)\*\*\*

CN 0 (Cytokines); 0 ( \*\*\*Interleukin\*\*\* -1); 0 ( \*\*\*Interleukin\*\*\* -2); 0 (Lipopolysaccharides); 0 (Phytohemagglutinins); 0 (Pokeweed Mitogens)

L10 ANSWER 18 OF 19 MEDLINE on STN

AN 1999314910 MEDLINE <<LOGINID::20080325>>

DN PubMed ID: 10438314

TI \*\*\*Interferon\*\*\* -gamma and \*\*\*interleukin\*\*\* -2 release by lymphocytes derived from the blood, mesenteric lymph nodes and intestines of normal sheep and those affected with \*\*\*paratuberculosis\*\*\* (Johne's disease).

AU Burrells C; Clarke C J; Colston A; Kay J M; Porter J; Little D; Sharp J M  
CS Moredun Research Institute, International Research Centre, Bush Loan, Penicuik, UK.. charles.burrells@cultor.com

SO Veterinary immunology and immunopathology, (1999 May) Vol. 68, No. 2-4, pp. 139-48.

Journal code: 8002006. ISSN: 0165-2427.

CY Netherlands

DT Journal; Article; (JOURNAL ARTICLE)  
(RESEARCH SUPPORT, NON-U.S. GOV'T)

LA English  
 FS Priority Journals; AIDS  
 EM 199908  
 ED Entered STN: 16 Aug 1999  
 Last Updated on STN: 16 Aug 1999  
 Entered Medline: 3 Aug 1999

AB This study sought to determine if T-cell cytokine responses to mycobacterial infections in sheep were similar to those in other species and if such responses correlated with prevailing gut pathology. Lymphocytes were isolated from the blood (PBL), mesenteric lymph nodes (MLN) and ileal lamina propria (LPL) of control sheep and of sheep with clinical Johne's disease due to infection with *Mycobacterium avium* ssp. *\*\*\*paratuberculosis\*\*\** (M.a. *\*\*\*paratuberculosis\*\*\**). These animals had previously been categorised into two groups exhibiting either the 'tuberculoid' (paucibacillary) form of lesion or the 'lepomatous' (multibacillary) form. Lymphocytes were examined for their capacity, following stimulation with johnin-PPD, to release *\*\*\*interferon\*\*\**-gamma (IFN-gamma) and *\*\*\*interleukin\*\*\** 2 (IL-2) characteristic of the Th1 subset of MHC Class II-restricted CD4+ (helper) T-cells in other species. The expression of the two cytokines appeared related to the type of histological lesion observed. Antigen-stimulated lymphocytes from the tuberculoid group exhibited greater release of IFN-gamma and IL-2 than lymphocytes from the lepomatous group suggesting a Th1-type of response in the former in which expression of IFN-gamma by PBL showed a significant positive correlation with that expressed by MLN and LPL. Lymphocytes from animals with lepomatous lesions released lesser mycobacterium-induced IFN-gamma and IL-2 indicating a diminished role for a Th1 subset in this group of sheep. Differences in cytokine expression were much more apparent with lymphocytes which were derived from MLN.

TI *\*\*\*Interferon\*\*\**-gamma and *\*\*\*interleukin\*\*\** -2 release by lymphocytes derived from the blood, mesenteric lymph nodes and intestines of normal sheep and those affected with *\*\*\*paratuberculosis\*\*\** (Johne's disease).

AB . . . lamina propria (LPL) of control sheep and of sheep with clinical Johne's disease due to infection with *Mycobacterium avium* ssp. *\*\*\*paratuberculosis\*\*\** (M.a. *\*\*\*paratuberculosis\*\*\**). These animals had previously been categorised into two groups exhibiting either the 'tuberculoid' (paucibacillary) form of lesion or the 'lepomatous' (multibacillary) form. Lymphocytes were examined for their capacity, following stimulation with johnin-PPD, to release *\*\*\*interferon\*\*\**-gamma (IFN-gamma) and *\*\*\*interleukin\*\*\** 2 (IL-2) characteristic of the Th1 subset of MHC Class II-restricted CD4+ (helper) T-cells in other species. The expression of. . .

CT Check Tags: Female  
 Animals  
*\*\*\* Enzyme-Linked Immunosorbent Assay: VE, veterinary\*\*\**  
 Ileum  
*\*\*\*\*Interferon Type II: BI, biosynthesis\*\*\**  
*\*\*\*\*Interleukin-2: BI, biosynthesis\*\*\**  
 Lymph Nodes: CY, cytology  
 \*Lymph Nodes: ME, metabolism  
 Lymphocyte Activation  
 Mesentery  
*\*\*\*\*Paratuberculosis: ME, metabolism\*\*\**  
 Sheep

\*Sheep Diseases: ME, metabolism  
 \*Th1 Cells: ME, metabolism  
 RN \*\*\*82115-62-6 (Interferon Type II)\*\*\*  
 CN 0 ( \*\*\*Interleukin\*\*\* -2)

L10 ANSWER 19 OF 19 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on  
 STN DUPLICATE 7

AN 1998:363991 BIOSIS <<LOGINID::20080325>>  
 DN PREV199800363991  
 TI \*\*\*Interferon\*\*\* -gamma and \*\*\*interleukin\*\*\* 4 gene expression in  
 cows infected with Mycobacterium \*\*\*paratuberculosis\*\*\* .  
 AU Sweeney, Raymond W. [Reprint author]; Jones, Douglas E.; Habecker, Perry;  
 Scott, Phillip  
 CS Dep. Clinical Studies-New Bolton Cent., Sch. Veterinary Med., Univ.  
 Pennsylvania, 382 W. Street Rd., Kennett Square, PA 19348, USA  
 SO American Journal of Veterinary Research, (July, 1998) Vol. 59, No. 7, pp.  
 842-847. print.  
 CODEN: AJVRAH. ISSN: 0002-9645.  
 DT Article  
 LA English  
 ED Entered STN: 27 Aug 1998  
 Last Updated on STN: 27 Aug 1998

AB Objective-To determine whether clinical progression of  
 \*\*\*paratuberculosis\*\*\* in cattle was associated with alterations in  
 cytokine gene expression in affected tissues. Animals-5 uninfected adult  
 Holstein cows, 7 adult Holstein cows naturally infected with Mycobacterium  
 \*\*\*paratuberculosis\*\*\* that did not have clinical signs of disease, and  
 4 adult Holstein cows naturally infected with M. \*\*\*paratuberculosis\*\*\*  
 that had progressive clinical signs of infection. Procedure-Samples of  
 ileum and cecal lymph nodes were obtained from each animal at the time of  
 slaughter. A reverse transcriptase-competitive polymerase chain reaction  
 \*\*\*assay\*\*\* was used to determine mRNA expression of \*\*\*interferon\*\*\*  
 -gamma (IFN-gamma) and \*\*\*interleukin\*\*\* 4 in each sample. Results-  
 \*\*\*Interferon\*\*\* -gamma gene expression was significantly higher in  
 ileum  
 and cecal lymph node samples from subclinically infected cows than from  
 clinically infected cows. Conclusions and Clinical Relevance-Progression  
 of \*\*\*paratuberculosis\*\*\* to clinical stages is associated with  
 reduced expression of IFN-gamma at site of infection. If immune response  
 to M. \*\*\*paratuberculosis\*\*\* can be manipulated so that IFN-gamma  
 expression is increased, resistance to infection in cattle might be  
 enhanced.

TI \*\*\*Interferon\*\*\* -gamma and \*\*\*interleukin\*\*\* 4 gene expression in  
 cows infected with Mycobacterium \*\*\*paratuberculosis\*\*\* .  
 AB Objective-To determine whether clinical progression of  
 \*\*\*paratuberculosis\*\*\* in cattle was associated with alterations in  
 cytokine gene expression in affected tissues. Animals-5 uninfected adult  
 Holstein cows, 7 adult Holstein cows naturally infected with Mycobacterium  
 \*\*\*paratuberculosis\*\*\* that did not have clinical signs of disease, and  
 4 adult Holstein cows naturally infected with M. \*\*\*paratuberculosis\*\*\*  
 that had progressive clinical signs of infection. Procedure-Samples of  
 ileum and cecal lymph nodes were obtained from each animal at the time of  
 slaughter. A reverse transcriptase-competitive polymerase chain reaction  
 \*\*\*assay\*\*\* was used to determine mRNA expression of \*\*\*interferon\*\*\*  
 -gamma (IFN-gamma) and \*\*\*interleukin\*\*\* 4 in each sample. Results-  
 \*\*\*Interferon\*\*\* -gamma gene expression was significantly higher in  
 ileum

and cecal lymph node samples from subclinically infected cows than from clinically infected cows. Conclusions and Clinical Relevance-Progression of \*\*\*paratuberculosis\*\*\* to clinical stages is associated with reduced expression of IFN-gamma at site of infection. If immune response to M. \*\*\*paratuberculosis\*\*\* can be manipulated so that IFN-gamma expression is increased, resistance to infection in cattle might be enhanced.

IT Major Concepts  
Immune System (Chemical Coordination and Homeostasis); Infection;  
Veterinary Medicine (Medical Sciences)

IT Diseases  
Mycobacterium- \*\*\*paratuberculosis\*\*\* infection: bacterial disease  
Mycobacterium Infections (MeSH)

IT Chemicals & Biochemicals  
\*\*\*interferon\*\*\* -gamma: gene expression; \*\*\*interleukin\*\*\* -4:  
gene expression

ORGN . . .  
Vertebrates, Nonhuman Mammals, Vertebrates

ORGN Classifier  
Mycobacteriaceae 08881  
Super Taxa  
Mycobacteria; Actinomycetes and Related Organisms; Eubacteria;  
Bacteria; Microorganisms  
Organism Name  
Mycobacterium- \*\*\*paratuberculosis\*\*\* : pathogen  
Taxa Notes  
Bacteria, Eubacteria, Microorganisms

=> s l10 and (interleukin 10)  
L11 7 L10 AND (INTERLEUKIN 10)

=> d 1-  
YOU HAVE REQUESTED DATA FROM 7 ANSWERS - CONTINUE? Y/(N):y

L11 ANSWER 1 OF 7 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
AN 2006:496945 BIOSIS <<LOGINID::20080325>>  
DN PREV200600503265  
TI Disturbed cytokine response to mycobacterium avium subspecies  
\*\*\*paratuberculosis\*\*\* is dysregulated in patients with Crohn's  
disease.  
AU Sibartie, Shomik; Keohane, John; Scully, Paul; O'Neill, Shaun; O'Mahony,  
Jim; O'Mahony, Liam; Shanahan, Fergus  
SO Gastroenterology, (APR 2006) Vol. 130, No. 4, Suppl. 2, pp. A240.  
Meeting Info.: Digestive Disease Week Meeting/107th Annual Meeting of the  
American-Gastroenterological-Association. Los Angeles, CA, USA. May 19  
-24, 2006. Amer Gastroenterol Assoc Inst.  
CODEN: GASTAB. ISSN: 0016-5085.  
DT Conference; (Meeting)  
Conference; Abstract; (Meeting Abstract)  
LA English  
ED Entered STN: 4 Oct 2006  
Last Updated on STN: 4 Oct 2006

L11 ANSWER 2 OF 7 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
AN 2004:178760 BIOSIS <<LOGINID::20080325>>  
DN PREV200400179647

TI Cytokine gene expression in peripheral blood mononuclear cells and tissues  
 of cattle infected with Mycobacterium avium subsp.  
 \*\*\*paratuberculosis\*\*\* : Evidence for an inherent proinflammatory gene  
 expression pattern.  
 AU Coussens, Paul M. [Reprint Author]; Verman, Nitin; Coussens, Marc A.;  
 Elftman, Michael D.; McNulty, Amanda M.  
 CS Department of Animal Science, Michigan State University, 1205H Anthony  
 Hall, East Lansing, MI, 48824, USA  
 coussens@msu.edu  
 SO Infection and Immunity, (March 2004) Vol. 72, No. 3, pp. 1409-1422. print.  
 ISSN: 0019-9567 (ISSN print).  
 DT Article  
 LA English  
 ED Entered STN: 31 Mar 2004  
 Last Updated on STN: 31 Mar 2004

L11 ANSWER 3 OF 7 CABA COPYRIGHT 2008 CABI on STN  
 AN 2006:49179 CABA <<LOGINID::20080325>>  
 DN 20063031176  
 TI Inflammatory cytokine gene expression in different types of granulomatous  
 lesions during asymptomatic stages of bovine \*\*\*paratuberculosis\*\*\*  
 AU Tanaka, S.; Sato, M.; Onitsuka, T.; Kamata, H.; Yokomizo, Y.  
 CS Comparative Pathology Section, Kyushu Research Station, National Institute  
 of Animal Health, Chuzan-cho 2702, Kagoshima 891-0105, Japan.  
 tanakas@affrc.go.jp  
 SO Veterinary Pathology, (2005) Vol. 42, No. 5, pp. 579-588. 41 ref.  
 Publisher: American College of Veterinary Pathologists Inc. Lawrence  
 ISSN: 0300-9858  
 DOI: 10.1354/vp.42-5-579  
 CY United States  
 DT Journal  
 LA English  
 ED Entered STN: 2 Mar 2006  
 Last Updated on STN: 2 Mar 2006

L11 ANSWER 4 OF 7 CAPLUS COPYRIGHT 2008 ACS on STN  
 AN 2007:906779 CAPLUS <<LOGINID::20080325>>  
 DN 147:275692  
 TI Sequences for Mycobacterium leprae-specific antigens, and methods for  
 treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in the early  
 stages and paucibacillary infections  
 IN Ottenhof, Tom Henricus Maria; Geluk, Annemieke; Pereira Sampaio, Elizabeth  
 PA Leiden University Medical Center, Neth.  
 SO PCT Int. Appl., 70pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	----	-----	-----	-----
PI	WO 2007091881	A2	20070816	WO 2006-NL50105	20060428
	WO 2007091881	A3	20071129		
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,			
		CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,			
		GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR,			
		KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX,			
		MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE,			



SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC,  
 VN, YU, ZA, ZM, ZW  
 RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,  
 IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ,  
 CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,  
 GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,  
 KG, KZ, MD, RU, TJ, TM, AP, EA, EP, OA  
 PRAI EP 2005-103576 A 20050429

L11 ANSWER 5 OF 7 CAPLUS COPYRIGHT 2008 ACS on STN  
 AN 2004:885718 CAPLUS <<LOGINID::20080325>>  
 DN 141:363746  
 TI Development of early-stage \*\*\*diagnostic\*\*\* method for Johne disease  
 by using anti-IL-10 antibody  
 AU Momotani, Eiichi; Mori, Yasuyuki  
 CS Natl. Agric. Bio-oriented Res. Org., Natl. Inst. Animal Health, Tsukuba,  
 305-0856, Japan  
 SO BRAIN Techno News (2004), 105, 18-24  
 CODEN: BTEEEC; ISSN: 1345-5958  
 PB Nogyo, Seibutsukei Tokutei Sangyo Gijutsu Kenkyu Kiko, Seibutsukei Tokutei  
 Sangyo Gijutsu Kenkyu Shien Senta  
 DT Journal; General Review  
 LA Japanese

L11 ANSWER 6 OF 7 CAPLUS COPYRIGHT 2008 ACS on STN  
 AN 2003:472526 CAPLUS <<LOGINID::20080325>>  
 DN 139:30816  
 TI Peptide T and analogs thereof for the stimulation of cytotoxic T  
 lymphocyte (CTL) responses and increasing secretion of \*\*\*interferon\*\*\*  
 .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use  
 IN Ruscetti, Francis W.; Ruff, Michael R.  
 PA The Government of the United States of America, as Represented by the  
 Secretary Department of Health and Human Services National Institutes of  
 Health, USA  
 SO PCT Int. Appl., 43 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2003050136	A2	20030619	WO 2002-US39109	20021206
	WO 2003050136	A3	20031204		
	W:				
	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,				
	CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,				
	GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,				
	LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,				
	PL, PT, RO, RU, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA,				
	UG, US, UZ, VN, YU, ZA, ZM, ZW				
	RW:				
	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,				
	KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,				
	FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ,				
	CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
	AU 2002357093	A1	20030623	AU 2002-357093	20021206
PRAI	US 2001-338971P	P	20011207		
	WO 2002-US39109	W	20021206		
OS	MARPAT 139:30816				

L11 ANSWER 7 OF 7 MEDLINE on STN  
 AN 2007416292 MEDLINE <<LOGINID::20080325>>  
 DN PubMed ID: 17502388  
 TI Influence of Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* on  
 colitis development and specific immune responses during disease.  
 AU Singh Udai P; Singh Shailesh; Singh Rajesh; Karls Russell K; Quinn  
 Frederick D; Potter Morris E; Lillard James W Jr  
 CS Brown Cancer Center, Department of Microbiology and Immunology, University  
 of Louisville, 580 S. Preston Street, Baxter II/Room 304C, Louisville, KY  
 40202, USA.  
 NC AI 57808 (United States NIAID)  
 GM 08248 (United States NIGMS)  
 MD 000525 (United States NCMHD)  
 RR 03034 (United States NCRR)  
 SO Infection and immunity, (2007 Aug) Vol. 75, No. 8, pp. 3722-8. Electronic  
 Publication: 2007-05-14.  
 Journal code: 0246127. ISSN: 0019-9567.  
 CY United States  
 DT Journal; Article; (JOURNAL ARTICLE)  
 (RESEARCH SUPPORT, N.I.H., EXTRAMURAL)  
 (RESEARCH SUPPORT, NON-U.S. GOV'T)  
 LA English  
 FS Priority Journals  
 EM 200709  
 ED Entered STN: 20 Jul 2007  
 Last Updated on STN: 7 Sep 2007  
 Entered Medline: 6 Sep 2007

=> s l11 and (anti-interleukin)  
 L12 0 L11 AND (ANTI-INTERLEUKIN)

=> s l11 and (antibody to interleukin)  
 L13 0 L11 AND (ANTIBODY TO INTERLEUKIN)

=> s l11 and ((antibody)2w(interleukin))  
 MISSING OPERATOR ANTIBODY)2W  
 The search profile that was entered contains terms or  
 nested terms that are not separated by a logical operator.

=> s l11 and (antibody(2w)interleukin)  
 L14 0 L11 AND (ANTIBODY(2W) INTERLEUKIN)

=> d kwic l11 1-  
 YOU HAVE REQUESTED DATA FROM 7 ANSWERS - CONTINUE? Y/(N):y

L11 ANSWER 1 OF 7 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
 TI Disturbed cytokine response to mycobacterium avium subspecies  
 \*\*\*paratuberculosis\*\*\* is dysregulated in patients with Crohn's  
 disease.  
 AB Background: Mycobacterium avium subspecies \*\*\*paratuberculosis\*\*\*  
 (MAP) has been a source of controversy since it was first suggested as a  
 possible cause for Crohn's disease. While. . . few studies have  
 examined the cellular immune response to MAP. Aim: To compare the  
 cellular response to Mycobacterium avium subspecies  
 \*\*\*paratuberculosis\*\*\* between Crohn's disease patients and healthy

volunteers. Methods: Peripheral blood mononuclear cells (PBMCs) were isolated from 24 Crohn's disease patients. . . .

IT . . . .  
system, blood and lymphatics, PBMC; phagocytic cell: immune system

IT Diseases  
Crohn's disease: digestive system disease, immune system disease, etiology, \*\*\*diagnosis\*\*\*

IT Chemicals & Biochemicals  
cytokines; IFN-gamma [ \*\*\*interferon\*\*\* -gamma]; IL-10 [ \*\*\*interleukin\*\*\* - \*\*\*10\*\*\* ]; TNF-alpha [tumor necrosis factor-alpha]; IL-6 [ \*\*\*interleukin\*\*\* -6]; IL-8 [ \*\*\*interleukin\*\*\* -8]; IL-2 [ \*\*\*interleukin\*\*\* -2]; IL-4 [ \*\*\*interleukin\*\*\* -4]

ORGN . . . .  
Primates, Vertebrates

ORGN Classifier  
Mycobacteriaceae 08881  
Super Taxa  
Mycobacteria; Actinomycetes and Related Organisms; Eubacteria; Bacteria; Microorganisms  
Organism Name  
Mycobacterium avium \*\*\*paratuberculosis\*\*\* (subspecies): pathogen  
Taxa Notes  
Bacteria, Eubacteria, Microorganisms

L11 ANSWER 2 OF 7 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN

TI Cytokine gene expression in peripheral blood mononuclear cells and tissues of cattle infected with Mycobacterium avium subsp.  
\*\*\*paratuberculosis\*\*\* : Evidence for an inherent proinflammatory gene expression pattern.

AB In cattle and other ruminants, infection with the intracellular pathogen Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* results in a granulomatous enteritis (Johne's disease) that is often fatal. The key features of host immunity to M. avium subsp. \*\*\*paratuberculosis\*\*\* infection include an appropriate early proinflammatory and cytotoxic response (Th1-like) that eventually gives way to a predominant antibody-based response (Th2-like).. . . this shift in the immune response occurs and the underlying molecular mechanisms involved is critical to future control measures and \*\*\*diagnosis\*\*\* . Previous studies have suggested that M. avium subsp. \*\*\*paratuberculosis\*\*\* may suppress gene expression in peripheral blood mononuclear cells (PBMCs) from infected cows, despite a continued inflammatory reaction at sites of infection. In the present study, we tested the hypothesis that exposure to M. avium subsp. \*\*\*paratuberculosis\*\*\* suppresses a proinflammatory gene expression pattern in PBMCs from infected cows. To do this, we examined expression of genes encoding \*\*\*interleukin\*\*\* -1alpha (IL-1alpha), IL-2, IL-4, IL-5, IL-6, IL-8, IL-10, IL-12p35, IL-16, and IL-18, as well as genes encoding gamma \*\*\*interferon\*\*\* (IFN-gamma), transforming growth factor beta (TGF-beta), and tumor necrosis factor alpha (TNF-alpha), in PBMCs, intestinal lesions, and mesenteric lymph nodes of cattle naturally infected with M. avium subsp.  
\*\*\*paratuberculosis\*\*\* . Cytokine gene expression in these cells and tissues was compared to expression in similar cells and tissues from control uninfected. . . IL-8, and IL-12p35, differential expression in PBMCs from infected and control cattle did not require stimulation with M. avium subsp. \*\*\*paratuberculosis\*\*\* . In fact, stimulation with M. avium subsp. \*\*\*paratuberculosis\*\*\* tended to reduce the differential

expression observed in infected and uninfected cows for genes encoding IFN-gamma, IL-1alpha, and IL-6. Only IL-10 gene expression was consistently enhanced by *M. avium* subsp. **\*\*\*paratuberculosis\*\*\*** stimulation of PBMCs from subclinically infected cattle. In ileal tissues from *M. avium* subsp. **\*\*\*paratuberculosis\*\*\*** -infected cattle, expression of the genes encoding IFN-gamma, TGF-beta, IL-5, and IL-8 was greater than the expression in comparable tissues from. . . was lower in tissues from infected cattle than in control tissues. Mesenteric lymph nodes draining sites of *M. avium* subsp. **\*\*\*paratuberculosis\*\*\*** infection expressed higher levels of IL-1alpha, IL-8, IL-2, and IL-10 mRNA than similar tissues from control uninfected cattle expressed. In. . . cattle. Taken together, our results suggest that cells or other mechanisms capable of limiting proinflammatory responses to *M. avium* subsp. **\*\*\*paratuberculosis\*\*\*** develop in infected cattle and that a likely place for development and expansion of these cell populations is the mesenteric. . .

IT . . .  
 lymph node: blood and lymphatics, digestive system, immune system;  
 peripheral blood mononuclear cell: blood and lymphatics, immune system

IT Diseases  
**\*\*\*paratuberculosis\*\*\*** : bacterial disease, infectious disease,  
 genetics, immunology, Johne's disease  
**\*\*\*Paratuberculosis\*\*\*** (MeSH)

IT Chemicals & Biochemicals  
 proinflammatory genes: expression pattern

ORGN . . .  
 Vertebrates

ORGN Classifier  
 Mycobacteriaceae 08881  
 Super Taxa  
 Mycobacteria; Actinomycetes and Related Organisms; Eubacteria;  
 Bacteria; Microorganisms  
 Organism Name  
 Mycobacterium avium ssp. **\*\*\*paratuberculosis\*\*\*** (subspecies):  
 pathogen  
 Taxa Notes  
 Bacteria, Eubacteria, Microorganisms

GEN cattle IFN-gamma gene [cattle **\*\*\*interferon\*\*\*** -gamma gene] (Bovidae);  
 cattle IL-1-alpha gene [cattle **\*\*\*interleukin\*\*\*** -1-alpha gene]  
 (Bovidae); cattle IL-10 gene [cattle **\*\*\*interleukin\*\*\*** - **\*\*\*10\*\*\***  
 gene] (Bovidae); cattle IL-12p35 gene [cattle **\*\*\*interleukin\*\*\*** -12p35  
 gene] (Bovidae); cattle IL-16 gene [cattle **\*\*\*interleukin\*\*\*** -16 gene]  
 (Bovidae); cattle IL-18 gene [cattle **\*\*\*interleukin\*\*\*** -18 gene]  
 (Bovidae); cattle IL-2 gene [cattle **\*\*\*interleukin\*\*\*** -2 gene]  
 (Bovidae); cattle IL-4 gene [cattle **\*\*\*interleukin\*\*\*** -4 gene]  
 (Bovidae); cattle IL-5 gene [cattle **\*\*\*interleukin\*\*\*** -5 gene]  
 (Bovidae); cattle IL-6 gene [cattle **\*\*\*interleukin\*\*\*** -6 gene]  
 (Bovidae); cattle IL-8 gene [cattle **\*\*\*interleukin\*\*\*** -8 gene]  
 (Bovidae); cattle TGF-beta gene [cattle transforming growth factor-beta  
 gene] (Bovidae); cattle TNF-alpha gene [cattle tumor necrosis factor-alpha  
 gene] (Bovidae)

L11 ANSWER 3 OF 7 CABA COPYRIGHT 2008 CABI on STN

TI Inflammatory cytokine gene expression in different types of granulomatous  
 lesions during asymptomatic stages of bovine **\*\*\*paratuberculosis\*\*\*** .

AB The granulomatous lesions in bovine **\*\*\*paratuberculosis\*\*\*** have been  
 classified into two types, i.e., the lepromatous type and the tuberculoid

type. To clarify the immunopathologic mechanisms at. . . the two types of lesions. Samples were obtained from noninfected control cows (n=5) and naturally infected cows (n=7) that were \*\*\*diagnosed\*\*\* by enzyme-linked immunosorbent \*\*\*assay\*\*\* (ELISA) and faecal culture test. Although none of the infected cows showed clinical signs, tuberculoid lesions were observed in five. . . and lepromatous lesions in two cows (lepromatous group). Among the cytokines examined by reverse transcription-polymerase chain reaction (RT-PCR), Th2-type cytokines \*\*\*interleukin\*\*\* -4 (IL-4) and IL-10, and Th1-type cytokine IL-2 were expressed more significantly in the lepromatous group than in the tuberculoid (P<0.01) and noninfected groups (P<0.05). No statistical differences were observed in the expression of \*\*\*interferon\*\*\* -gamma, IL-1 beta, TNF-alpha, and GM-CSF among lepromatous, tuberculoid, and noninfected groups. Expression of proinflammatory cytokine IL-12 mRNA, however, did not. . . influenced by alterations in Th1/Th2-type cytokine production and that IL-18 may play an important role in a Th1-to-Th2 switch in \*\*\*paratuberculosis\*\*\* .

CT cows; cytokines; disease course; gene expression; genes; granuloma; histopathology; immunopathology; \*\*\*interferon\*\*\* ; \*\*\*interleukin\*\*\* 1; \*\*\*interleukin\*\*\* \*\*\*10\*\*\* ; \*\*\*interleukin\*\*\* 2; \*\*\*interleukin\*\*\* 4; \*\*\*interleukins\*\*\* ; messenger RNA; \*\*\*paratuberculosis\*\*\* ; tumour necrosis factor

ST \*\*\*interleukin\*\*\* 18

ORGN cattle; Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\*

L11 ANSWER 4 OF 7 CAPLUS COPYRIGHT 2008 ACS on STN

TI Sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in the early stages and paucibacillary infections

AB The current invention discloses new Mycobacterium leprae antigens to be used in methods and means for detection and \*\*\*diagnostics\*\*\* of M. leprae infections in subjects, in particular in the early stages of infection and in paucibacillary infections, which remain undetected using conventional \*\*\*diagnostic\*\*\* methods. The antigens disclosed in the invention are specific for M. leprae and the \*\*\*diagnostic\*\*\* method does not yield 'false pos.' results in individuals having an immune response against other Mycobacterial species, such as M. tuberculosis, M. bovis, M. \*\*\*paratuberculosis\*\*\* , M. avium, M. smegmatis,, M. ulcerans, M. microti, and M. marinum, or BCG vaccinated individuals. Thus, using bioinformatic anal. the. . . in human leprosy tissue. Paucibacillary and reactional leprosy patients and healthy household contacts of leprosy patients produced significant levels of \*\*\*interferon\*\*\* (IFN)-.gamma. in response to the five unique M. leprae antigens encoded by ML0576, ML1989, ML1990, ML2283 and ML2567. Provided are. . .

ST sequence Mycobacterium leprae antigen epitope \*\*\*diagnoses\*\*\* infection; leprosy immunodiagnosis Mycobacterium leprae antigen epitope; vaccine Mycobacterium leprae antigen epitope

IT Receptors

RL: BSU (Biological study, unclassified); BIOL (Biological study) (4-1BB, anti-4-1BB agonistic antibody as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Human groups (Brazilian patients; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae,

particularly in early stages and paucibacillary infections)

IT Genetic element  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (CpG island, CpG, as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Histocompatibility antigens  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (HLA, class I, identifying T-cell epitopes for, using computer algorithms; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Histocompatibility antigens  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (HLA, class II, identifying T-cell epitopes for, using computer algorithms; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Proteins  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (LAG3 (lymphocyte activation gene-3), sol., as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML0573, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML0574, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML0575, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML0576, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Antigens  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL

(Biological study); USES (Uses)  
 (ML0576; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1602, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1603, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1604, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1788, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1989, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Antigens  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (ML1989; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML1990, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Antigens  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use);

PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (ML1990; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML2283, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Antigens  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (ML2283; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Gene, microbial  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (ML2567, expressed in human leprosy tissue; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Antigens  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (ML2567; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Lipopeptides  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (Pam3Cys, as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Immunostimulants  
 (adjuvants, DA/TDB; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Immunostimulants  
 (adjuvants, DDA/MPL; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Immunostimulants  
 (adjuvants; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Monocyte  
 (anal., in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)



IT     \*\*\*Diagnostic\*\*\*     agents  
       Vaccines  
          (antigens or epitopes as; sequences for Mycobacterium leprae-specific  
          antigens, and methods for treating and     \*\*\*diagnosing\*\*\*     M. leprae,  
          particularly in early stages and paucibacillary infections)

IT     Lipid A  
       Lipopolysaccharides  
       RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
          (as adjuvant; sequences for Mycobacterium leprae-specific antigens, and  
          methods for treating and     \*\*\*diagnosing\*\*\*     M. leprae, particularly  
          in early stages and paucibacillary infections)

IT     Mycobacterium  
       (as recombinant expression host; sequences for Mycobacterium  
       leprae-specific antigens, and methods for treating and  
       \*\*\*diagnosing\*\*\*     M. leprae, particularly in early stages and  
       paucibacillary infections)

IT     Flagellins  
       RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
          (bacterial, as adjuvant; sequences for Mycobacterium leprae-specific  
          antigens, and methods for treating and     \*\*\*diagnosing\*\*\*     M. leprae,  
          particularly in early stages and paucibacillary infections)

IT     CD40 (antigen)  
       RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
          (binding CD40 ligand or antibody, as adjuvant; sequences for  
          Mycobacterium leprae-specific antigens, and methods for treating and  
          \*\*\*diagnosing\*\*\*     M. leprae, particularly in early stages and  
          paucibacillary infections)

IT     Mammalia  
       (     \*\*\*diagnosis\*\*\*     and therapy; sequences for Mycobacterium  
       leprae-specific antigens, and methods for treating and  
       \*\*\*diagnosing\*\*\*     M. leprae, particularly in early stages and  
       paucibacillary infections)

IT     Mycobacterium avium  
       Mycobacterium bovis  
       Mycobacterium marinum  
       Mycobacterium microti  
       Mycobacterium smegmatis  
       Mycobacterium tuberculosis  
       Mycobacterium ulcerans  
          (differentiating from; sequences for Mycobacterium leprae-specific  
          antigens, and methods for treating and     \*\*\*diagnosing\*\*\*     M. leprae,  
          particularly in early stages and paucibacillary infections)

IT     Leprosy  
       (early stages     \*\*\*diagnosis\*\*\*     ; sequences for Mycobacterium  
       leprae-specific antigens, and methods for treating and  
       \*\*\*diagnosing\*\*\*     M. leprae, particularly in early stages and  
       paucibacillary infections)

IT     T cell (lymphocyte)  
       (epitopes; sequences for Mycobacterium leprae-specific antigens, and  
       methods for treating and     \*\*\*diagnosing\*\*\*     M. leprae, particularly  
       in early stages and paucibacillary infections)

IT     Epitopes  
       (from ML0576, ML1989, ML1990, ML2283 and ML2567 antigens; sequences for  
       Mycobacterium leprae-specific antigens, and methods for treating and  
       \*\*\*diagnosing\*\*\*     M. leprae, particularly in early stages and  
       paucibacillary infections)

IT     T cell (lymphocyte)

(helper cell, measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Algorithm  
(identifying HLA class I and/or class II T-cell epitopes using; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Diagnosis\*\*\*  
(immunodiagnosis, of ML0576, ML1989, ML1990, ML2283 and ML2567 antigens; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Blood analysis  
(in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Interleukin\*\*\* \*\*\*10\*\*\*  
     \*\*\*Interleukin\*\*\* 15  
     \*\*\*Interleukin\*\*\* 2  
     \*\*\*Interleukin\*\*\* 4  
     \*\*\*Interleukin\*\*\* 6

Macrophage inflammatory protein 1.beta.  
 Transforming growth factor .beta.  
 Tumor necrosis factors

RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
 (measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Antibodies and Immunoglobulins  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (monoclonal, anti-4-1BB, agonistic, as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Genome  
(of M. leprae, identifying unique antigen gene candidates in; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Protein sequences  
(of M. leprae-specific antigens ML0576, ML1989, ML1990, ML2283 and ML2567; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT DNA sequences  
(of M. leprae-specific genes ML0576, ML1989, ML1990, ML2283 and ML2567; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Blood cell  
(of infected subject, IFN-.gamma. response in; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and

paucibacillary infections)

IT \*\*\*Interleukin\*\*\* 12  
 RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
 (p70, measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Human  
 (patients; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Infection  
 (paucibacillary, \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Bioinformatics  
 (sequence annotation, M. leprae unique genes; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Molecular cloning  
 Mycobacterium leprae  
 Test kits  
 (sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Skin  
 (test, by applying antigen under top skin; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT Mycobacterium BCG  
 (vaccine, differentiating from; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Interferons\*\*\*  
 RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
 (.alpha., measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Interferons\*\*\*  
 RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
 (.beta., measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT \*\*\*Interferons\*\*\*  
 RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
 (.gamma., measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and

\*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 141256-04-4, QS21  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (MPL, as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 946442-88-2 946442-91-7  
 RL: PRP (Properties)  
 (Unclaimed; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in the early stages and paucibacillary infections)

IT 946400-78-8 946400-79-9 946400-80-2 946400-81-3 946400-82-4  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (amino acid sequence, epitope; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 946442-52-0 946442-53-1 946442-54-2 946442-55-3 946442-56-4  
 RL: ADV (Adverse effect, including toxicity); ARU (Analytical role, unclassified); BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (amino acid sequence; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 24939-03-5, Poly(I:C) 87420-41-5, Pam3Cys 911642-39-2, IC 31  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (as adjuvant; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 83869-56-1, GM-CSF  
 RL: ARU (Analytical role, unclassified); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)  
 (measuring response, in \*\*\*diagnosis\*\*\* ; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 946442-57-5, DNA (Mycobacterium leprae gene ML0576) 946442-58-6, DNA (Mycobacterium leprae gene ML1989) 946442-59-7, DNA (Mycobacterium leprae gene ML1990) 946442-60-0, DNA (Mycobacterium leprae gene ML2283) 946442-61-1, DNA (Mycobacterium leprae gene ML2567)  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (nucleotide sequence; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in early stages and paucibacillary infections)

IT 946442-98-4 946442-99-5 946443-00-1 946443-01-2 946443-02-3  
 946443-03-4 946443-04-5 946443-05-6 946443-06-7 946443-07-8  
 946443-08-9 946443-09-0  
 RL: PRP (Properties)  
 (unclaimed nucleotide sequence; sequences for Mycobacterium leprae-specific antigens, and methods for treating and \*\*\*diagnosing\*\*\* M. leprae, particularly in the early stages and

paucibacillary infections)

IT 946442-86-0 946442-87-1 946442-89-3 946442-90-6 946442-92-8  
946442-93-9 946442-94-0 946442-95-1 946442-96-2 946442-97-3  
RL: PRP (Properties)  
(unclaimed protein sequence; sequences for Mycobacterium  
leprae-specific antigens, and methods for treating and  
\*\*\*diagnosing\*\*\* M. leprae, particularly in the early stages and  
paucibacillary infections)

L11 ANSWER 5 OF 7 CAPLUS COPYRIGHT 2008 ACS on STN

TI Development of early-stage \*\*\*diagnostic\*\*\* method for Johne disease  
by using anti-IL-10 antibody

AB A review on early-stage \*\*\*diagnosis\*\*\* of Johne's disease (  
\*\*\*paratuberculosis\*\*\* ) in cattle by modified \*\*\*interferon\*\*\*  
.gamma. ELISA \*\*\*assay\*\*\* using IL-10 neutralizing antibody, and its  
effectiveness.

ST review cattle Johne disease \*\*\*diagnosis\*\*\* ELISA \*\*\*interleukin\*\*\*  
\*\*\*10\*\*\* antibody; \*\*\*paratuberculosis\*\*\* cattle \*\*\*diagnosis\*\*\*  
\*\*\*interferon\*\*\* gamma ELISA review

IT Bos taurus  
Mycobacterium avium \*\*\*paratuberculosis\*\*\*  
(early-stage \*\*\*diagnosis\*\*\* method for Johne's disease using  
anti-IL-10 antibody)

IT \*\*\*Interleukin\*\*\* \*\*\*10\*\*\*  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(early-stage \*\*\*diagnosis\*\*\* method for Johne's disease using  
anti-IL-10 antibody)

IT Immunoassay  
(enzyme-linked immunosorbent \*\*\*assay\*\*\* ; early-stage  
\*\*\*diagnosis\*\*\* method for Johne's disease using anti-IL-10  
antibody)

IT \*\*\*Diagnosis\*\*\*  
(immunodiagnosis; early-stage \*\*\*diagnosis\*\*\* method for Johne's  
disease using anti-IL-10 antibody)

IT Infection  
( \*\*\*paratuberculosis\*\*\* , Johne's disease; early-stage  
\*\*\*diagnosis\*\*\* method for Johne's disease using anti-IL-10  
antibody)

IT Antibodies and Immunoglobulins  
RL: ARU (Analytical role, unclassified); BSU (Biological study,  
unclassified); DGN (Diagnostic use); ANST (Analytical study); BIOL  
(Biological study); USES (Uses)  
(to IL-10; early-stage \*\*\*diagnosis\*\*\* method for Johne's disease  
using anti-IL-10 antibody)

IT \*\*\*Interferons\*\*\*  
RL: ANT (Analyte); BSU (Biological study, unclassified); DGN (Diagnostic  
use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
(.gamma.; early-stage \*\*\*diagnosis\*\*\* method for Johne's disease  
using anti-IL-10 antibody)

L11 ANSWER 6 OF 7 CAPLUS COPYRIGHT 2008 ACS on STN

TI Peptide T and analogs thereof for the stimulation of cytotoxic T  
lymphocyte (CTL) responses and increasing secretion of \*\*\*interferon\*\*\*  
.gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use

AB . . . in a subject, comprising administering a CTL activity-stimulating  
amt. of peptide T or an analog thereof. A method of increasing .gamma.-  
\*\*\*interferon\*\*\* (IFN-.gamma.) secretion in a subject comprises

administering an IFN-.gamma. secretion-increasing amt. of peptide T or an analog thereof. A method of increasing \*\*\*interleukin\*\*\* 2 (IL-2) secretion in a subject comprises administering an IL-2 secretion-increasing amt. of peptide T or an analog thereof. A method of treating a subject \*\*\*diagnosed\*\*\* as having a disease assocd. with reduced CTL activity comprises administering a CTL activity-stimulating amt. of peptide T or an analog thereof. A method of treating a subject \*\*\*diagnosed\*\*\* as having a disease assocd. with reduced IFN-.gamma. activity comprises administering an IFN-.gamma. activity-stimulating amt. of peptide T or an analog thereof. A method of treating a subject \*\*\*diagnosed\*\*\* as having a disease assocd. with reduced IL-2 activity comprises administering an IL-2 activity-stimulating amt. of peptide T or an. . .

- ST peptide T analog cytotoxic T lymphocyte response stimulation;  
     \*\*\*interferon\*\*\* gamma secretion peptide T;   \*\*\*interleukin\*\*\* 2  
     secretion peptide T
- IT Lymphoma  
     (B-cell; peptide T and analogs for stimulation of cytotoxic T  
     lymphocyte responses and increasing secretion of   \*\*\*interferon\*\*\*  
     .gamma. and   \*\*\*interleukin\*\*\* 2, and therapeutic use)
- IT Lymphoma  
     (T-cell; peptide T and analogs for stimulation of cytotoxic T  
     lymphocyte responses and increasing secretion of   \*\*\*interferon\*\*\*  
     .gamma. and   \*\*\*interleukin\*\*\* 2, and therapeutic use)
- IT Tumor necrosis factors  
     RL: BSU (Biological study, unclassified); BIOL (Biological study)  
     (TNF-.alpha.; peptide T and analogs for stimulation of cytotoxic T  
     lymphocyte responses and increasing secretion of   \*\*\*interferon\*\*\*  
     .gamma. and   \*\*\*interleukin\*\*\* 2, and therapeutic use)
- IT Carcinoma  
     (adenocarcinoma; peptide T and analogs for stimulation of cytotoxic T  
     lymphocyte responses and increasing secretion of   \*\*\*interferon\*\*\*  
     .gamma. and   \*\*\*interleukin\*\*\* 2, and therapeutic use)
- IT AIDS (disease)  
     (and AIDS-related lymphoma or sarcoma; peptide T and analogs for  
     stimulation of cytotoxic T lymphocyte responses and increasing  
     secretion of   \*\*\*interferon\*\*\* .gamma. and   \*\*\*interleukin\*\*\* 2,  
     and therapeutic use)
- IT Infection  
     (bacterial; peptide T and analogs for stimulation of cytotoxic T  
     lymphocyte responses and increasing secretion of   \*\*\*interferon\*\*\*  
     .gamma. and   \*\*\*interleukin\*\*\* 2, and therapeutic use)
- IT Neoplasm  
     (blastoma; peptide T and analogs for stimulation of cytotoxic T  
     lymphocyte responses and increasing secretion of   \*\*\*interferon\*\*\*  
     .gamma. and   \*\*\*interleukin\*\*\* 2, and therapeutic use)
- IT Urogenital system  
     (cancer; peptide T and analogs for stimulation of cytotoxic T  
     lymphocyte responses and increasing secretion of   \*\*\*interferon\*\*\*  
     .gamma. and   \*\*\*interleukin\*\*\* 2, and therapeutic use)
- IT Esophagus, neoplasm  
     Head and Neck, neoplasm  
     Head and Neck, neoplasm  
     (carcinoma; peptide T and analogs for stimulation of cytotoxic T  
     lymphocyte responses and increasing secretion of   \*\*\*interferon\*\*\*  
     .gamma. and   \*\*\*interleukin\*\*\* 2, and therapeutic use)
- IT Uterus, neoplasm

(cervix, carcinoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
 Uterus, neoplasm  
 (cervix; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Intestine, neoplasm  
 (colon; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Intestine, neoplasm  
 (colorectal; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT T cell (lymphocyte)  
 (cytotoxic; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
 (esophageal; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Mycosis  
 (fungoides; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Neuroglia, neoplasm  
 (glioblastoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
 Carcinoma  
 (head and neck squamous cell carcinoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
 Carcinoma  
 (head and neck; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Neoplasm  
 (histiocytoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Hypoxia  
 (hypoxic tumors; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Fungi  
 Parasite  
 (infection; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma

(laryngeal squamous cell; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Neoplasm  
(metastasis; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Skin, neoplasm  
(mycosis fungoides; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Histiocyte  
(neoplasm, histiocytoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Nerve, neoplasm  
(neuroblastoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Lymphoma  
(non-Hodgkin's; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
(oral squamous cell; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Actinobacillus pleuropneumoniae

Adenoma

Alternaria alternata

Anti-AIDS agents

Antibacterial agents

Antimalarials

Antitumor agents

Antiviral agents

Aspergillus fumigatus

Bacillus anthracis

Bladder, neoplasm

Blastomyces dermatitidis

Brain, neoplasm

Brucella

Brucella melitensis

CD8-positive T cell

Campylobacter

Candida albicans

Carcinoma

Carcinoma

Chlamydia pneumoniae

Chlamydia trachomatis

Chlamydophila psittaci

Clostridium

Clostridium tetani

Coccidioides immitis



Coronavirus  
Coxiella burnetii  
Cryptococcus neoformans  
Dengue virus  
Eastern equine encephalitis virus  
Ebola virus  
Ehrlichia  
Ehrlichia ruminantium  
Entamoeba histolytica  
Escherichia coli  
Fungicides  
Haemophilus  
Haemophilus ducreyi  
Haemophilus influenzae  
Hantavirus  
Hematopoietic neoplasm  
Hepatitis A virus  
Hepatitis B virus  
Hepatitis C virus  
Hepatitis E virus  
Hepatitis delta virus  
Histoplasma capsulatum  
Hodgkin's disease  
Human  
Human T-lymphotropic virus 1  
Human adenovirus  
Human coxsackievirus  
Human immunodeficiency virus  
Human immunodeficiency virus 1  
Human immunodeficiency virus 2  
Human papillomavirus  
Human poliovirus  
Immunostimulants  
Influenza A virus  
Influenza B virus  
Japanese encephalitis virus  
Kidney, neoplasm  
Lassa virus  
Legionella  
Legionella pneumophila  
Leishmania  
Leishmania major  
Leukemia  
Listeria ivanovii  
Listeria monocytogenes  
Liver, neoplasm  
Lung, neoplasm  
Lymphoma  
Mammary gland, neoplasm  
Mannheimia haemolytica  
Marburg virus  
Measles virus  
Melanoma  
Multiple myeloma  
Mumps virus  
Murray Valley encephalitis virus  
Mycobacterium BCG

Mycobacterium africanum  
Mycobacterium avium  
Mycobacterium avium \*\*\*paratuberculosis\*\*\*  
Mycobacterium bovis  
Mycobacterium intracellulare  
Mycobacterium kansasii  
Mycobacterium marinum  
Mycobacterium tuberculosis  
Mycobacterium ulcerans  
Myeloid leukemia  
Neisseria gonorrhoeae  
Neisseria meningitidis  
Neoplasm  
Nervous system, neoplasm  
Neuroglia, neoplasm  
Nocardia  
Nocardia asteroides  
Ovary, neoplasm  
Pancreas, neoplasm  
Paracoccidioides brasiliensis  
Parasiticides  
Pasteurella  
Pasteurella multocida  
Penicillium marneffeii  
Plasmodium (malarial genus)  
Plasmodium falciparum  
Plasmodium malariae  
Plasmodium vivax  
Pneumocystis carinii  
Polyomavirus  
Prostate gland, neoplasm  
Pseudomonas  
Pseudomonas aeruginosa  
Rabies virus  
Respiratory syncytial virus  
Rhinovirus  
Rickettsia  
Rift Valley fever virus  
Rotavirus A  
Rotavirus B  
Rotavirus C  
Rous sarcoma virus  
Rubella virus  
Salmonella  
Salmonella typhi  
Sarcoma  
Schistosoma  
Schistosoma mansoni  
Shigella  
Simian immunodeficiency virus  
Sindbis virus  
Skin, neoplasm  
St. Louis encephalitis virus  
Staphylococcus aureus  
Staphylococcus epidermidis  
Streptococcus agalactiae  
Streptococcus pyogenes

Testis, neoplasm  
 Toxoplasma gondii  
 Trypanosoma brucei  
 Trypanosoma cruzi  
 Variola virus  
 Vesicular stomatitis virus  
 Vibrio cholerae  
 West Nile virus  
 Yellow fever virus  
 Yersinia  
 Yersinia enterocolitica  
 Yersinia pestis  
 (peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Cytokines  
     \*\*\*Interleukin\*\*\* 1  
     \*\*\*Interleukin\*\*\* \*\*\*10\*\*\*  
     \*\*\*Interleukin\*\*\* 12  
     \*\*\*Interleukin\*\*\* 13  
     \*\*\*Interleukin\*\*\* 2  
     \*\*\*Interleukin\*\*\* 4  
     \*\*\*Interleukin\*\*\* 6  
     \*\*\*Interleukin\*\*\* 8  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Peptides, biological studies  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
 (pulmonary squamous cell; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Neoplasm  
 (solid, carcinoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Head and Neck, neoplasm  
 Head and Neck, neoplasm  
 Larynx, neoplasm  
 Lung, neoplasm  
 Mouth, neoplasm  
 (squamous cell carcinoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Carcinoma  
 (squamous cell; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\*

.gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Pharynx, neoplasm  
(throat squamous cell carcinoma; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT Infection  
(viral; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT \*\*\*Interferons\*\*\*  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(.gamma.; peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT 106362-32-7D, C-terminal derivs.  
RL: PAC (Pharmacological activity); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT 106362-32-7, Peptide T 106362-32-7D, Peptide T, analogs  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(peptide T and analogs for stimulation of cytotoxic T lymphocyte responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

IT 107531-09-9 107531-11-3 107531-12-4 107531-14-6 118936-25-7  
118936-26-8 118936-27-9 118936-30-4 118936-31-5 118936-32-6  
118957-86-1 119386-95-7  
RL: PRP (Properties)  
(unclaimed sequence; peptide T and analogs thereof for the stimulation of cytotoxic T lymphocyte (CTL) responses and increasing secretion of \*\*\*interferon\*\*\* .gamma. and \*\*\*interleukin\*\*\* 2, and therapeutic use)

L11 ANSWER 7 OF 7 MEDLINE on STN

TI Influence of Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* on colitis development and specific immune responses during disease.

AB . . . and intramural inflammation observed in cases of inflammatory bowel diseases (IBD) and veterinary Johne's disease suggests that Mycobacterium avium subsp. \*\*\*paratuberculosis\*\*\* is a causative agent. However, an incomplete understanding of the immunological steps responsible for the pathologies of IBD makes this conclusion uncertain. Sera from \*\*\*interleukin\*\*\* - \*\*\*10\*\*\* -deficient (IL-10(-/-)) mice with spontaneous colitis displayed significantly higher M. avium subsp. \*\*\*paratuberculosis\*\*\* -specific immunoglobulin G2a antibody responses than did sera from similar mice without disease. Pathogen-free IL-10(-/-) mice received control vehicle or the vehicle containing heat-killed or live M. avium subsp. \*\*\*paratuberculosis\*\*\* . Mucosal CD4(+) T cells from the mice that developed colitis proliferated and secreted higher levels of gamma \*\*\*interferon\*\*\* and tumor necrosis factor alpha after ex vivo stimulation with a Vbeta11(+) T-cell receptor-restricted peptide from the MPT59 antigen (Ag85B). . . .

CT . . .

Antigens, Bacterial: IM, immunology  
     CD4-Positive T-Lymphocytes: IM, immunology  
     Colitis: IM, immunology  
 \*Colitis: MI, microbiology  
 \*Colitis: PA, pathology  
     Disease Models, Animal  
         \*\*\* Enzyme-Linked Immunosorbent Assay\*\*\*  
     Humans  
     Immunoglobulin G: BL, blood  
         \*\*\* Interferon Type II: BI, biosynthesis\*\*\*  
         \*\*\* Interleukin-10: DF, deficiency\*\*\*  
     Intestinal Mucosa: IM, immunology  
     Ligands  
     Mice  
     Mice, Knockout  
         \*\*\*\*Mycobacterium avium subsp. paratuberculosis: IM, immunology\*\*\*  
         \*\*\*\*Paratuberculosis: IM, immunology\*\*\*  
         \*\*\*\*Paratuberculosis: PA, pathology\*\*\*  
     Peptides: IM, immunology  
     Receptors, Antigen, T-Cell: IM, immunology  
     Receptors, CXCR3  
     Receptors, Chemokine: AG, agonists  
     Receptors, Chemokine: IM, immunology  
 . . .  
 RN       \*\*\*130068-27-8 (Interleukin-10)\*\*\* ;     \*\*\*82115-62-6 (Interferon Type\*\*\*  
       \*\*\*       II)\*\*\*  
 CN.     . . . 0 (Peptides); 0 (Receptors, Antigen, T-Cell); 0 (Receptors, CXCR3);  
           0 (Receptors, Chemokine); 0 (Tumor Necrosis Factor-alpha); 0 (antigen 85B,  
           Mycobacterium     \*\*\*paratuberculosis\*\*\* )